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PATENT APPLICATION TRANSMITTAL LETTER

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To the Commissioner of Patents and Trademarks:

Transmitted herewith for filing under 35 U.S.C. 111 and 37 CFR 1.53 is the patent application of:

Wayne A. Noda, et al.

entitled:

INDWELLING HEAT EXCHANGE CATHETER AND METHOD OF USING SAME

Enclosed are:

- ☒ 40 pages of written description, claims and abstract.
- ☒ 21 sheets of informal drawings.
- ☒ executed declaration and power of attorney by the inventors.
- ☒ a verified statement to establish small entity status under 37 CFR 1.9 and 1.27.
- ☒ return postcard.

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CLAIMS AS FILED

	NUMBER FILED	NUMBER EXTRA	RATE	FEE
BASIC FEE			\$ 790	\$ 790
TOTAL CLAIMS	63 - 20 =	* 43	X \$ 22	\$946
INDEPENDENT CLAIMS	8 - 3 =	* 5	X \$ 82	\$410
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*NUMBER EXTRA MUST BE ZERO OR LARGER			TOTAL	\$2,146
If applicant has small entity status under 37 CFR 1.9 and 1.27, then divide total fee by 2 and enter amount here.			SMALL ENTITY TOTAL	\$1,073

- ☒ A check in the amount of \$ 1,073.00 to cover the filing fee is enclosed.
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1. Patent Application transmittal in duplicate;
2. Patent Application in 40 pages, 63 claims;
3. Twenty-one (21) sheets of informal drawings;
4. Check in the amount of \$1,073.00 for filing fee;
5. Signed Declaration and Power of Attorney by Inventors;
6. Small Entity Statement;
7. Return receipt postcard.

Each of the above-identified documents are enclosed herewith.

Respectfully submitted,



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INDWELLING HEAT EXCHANGE CATHETER AND METHOD OF USING SAME

Background of the Invention

Field of the Invention

This invention relates generally to apparatus and methods for producing heat exchange with body tissue, and more specifically to methods and apparatus for the hypothermic treatment of a body fluid in a body conduit.

Discussion of the Prior Art

Many of the advantages of hypothermia are well known. By way of example, it has been found particularly desirable to lower the temperature of body tissue in order to reduce the metabolism of the body. In stroke, trauma and several other pathological conditions, hypothermia also reduces the permeability of the blood/brain barrier. It inhibits release of damaging neurotransmitters and also inhibits calcium-mediated effects. Hypothermia inhibits brain edema and lowers intracranial pressure.

In the past, hypothermic treatment has been typically addressed systemically, meaning that the overall temperature of the entire body has been lowered to achieve the advantages noted above. This has been particularly desirable in surgical applications where the reduced metabolism has made it possible to more easily accommodate lengthy operative procedures. An example of this systemic approach includes catheters for transferring heat to or from blood flowing within a patient's vessel, as disclosed by Ginsburg in U.S. Patent No. 5,486,208. A closed loop heat exchange catheter is also disclosed by Saab in U.S. Patent No. 5,624,392.

A cooling device for whole-body hyperthermia that utilizes the circulatory system of the body is known to be more efficient since the entire volume of the body is constantly perfused with the cold fluid at a capillary level.

Likewise, various other means of cooling the body have been tried with cooling blankets, ice water bladder lavages, ice baths, esophageal catheters and their associated methods. All of these devices require a considerable time to cool the body since the primary heat transfer occurs through the skin or the skull. A more efficient body cooling device that can quickly cool and accurately control the body temperature is required.

Summary of the Invention

A heat exchange catheter and method of operation are included in the present invention. The method is adapted to produce hypothermia or hyperthermia, typically in a selected portion of the body without substantially varying the temperature of the remaining portions of the body. The selected body portion will usually be associated with a body conduit which conveys a body fluid to the selected body portion. Of particular interest are the organs of the body which are commonly nourished and maintained by a flow of blood in the arterial system. For example, a flow of blood is introduced to the brain through the carotid artery. Of course the temperature of this blood is usually at the normal body temperature.

By positioning a heat exchange catheter in the body conduit, heat can be added to or removed from the body fluid to heat or cool the selected body portion. For example, the heat exchange catheter can be disposed in the carotid artery where the arterial blood flowing to the brain can be cooled. The flow of cooled blood to the brain reduces the temperature of the brain thereby resulting in cerebral hypothermia. Importantly, this temperature reduction occurs primarily and selectively in the brain; the remaining portions of the body maintain a generally normal body temperature. In accordance with this method, the selected body

portion, such as the brain, can be cooled thereby providing the advantages associated with hypothermia for this body portion. The remainder of the body, such as the portions other than the brain, do not experience the reduction in temperature. Furthermore, the invention is intended to remotely alter temperature in a region other than the point of introduction into the body. This is different than devices intended for systemic temperature control.

Several factors are of interest in effecting heat transfer in a heat exchanger. These factors include, for example, the convection heat transfer coefficient of the two fluids involved in the heat exchange, as well as the thermal conductivity and thickness of the barrier between the two fluids. Other factors include the relative temperature differential between the fluids, as well as the contact area and residence time of heat transfer. The Reynolds number for each fluid stream affects boundary layers, turbulence and laminar flow.

Notwithstanding the need for localized hypothermia, there will always be those procedures which call for systemic hypothermia. Many of the advantages associated with the present invention will greatly facilitate those procedures, for example, by decreasing the number and complexity of operative steps, increasing the heat transfer capacity of the device, and addressing other concerns such as the formation of blood clots.

In one aspect of the invention a catheter is provided with an elongate configuration, a proximal end and a distal end. An outer tube having a first lumen extends between the distal end and proximal end of the catheter, and an inner tube having a second lumen is disposed within the first lumen of the outer tube. Portions of the inner tube define a first flow path extending along the second lumen, while portions of the tubes define a second flow path extending between the first tube and the second tube. A plurality of hollow fibers provide fluid communication between the first and second flow paths, and a heat exchange fluid is disposed in the hollow fibers to cool the fibers.

In another aspect of the invention, a method for making a heat exchange catheter includes the steps of providing first and second tubes having first and second lumens, respectively. A plurality of hollow fibers are connected between a first flow path extending

along the second lumen and a second flow path extending along the first lumen outwardly of the second tube. The method further comprises the step of insuring that the second tube is axially or rotationally movable relative to the first tube in order to vary the configuration of the hollow fibers.

In a further aspect of the invention, a method for operating a heat exchange catheter includes the steps of inserting into a body conduit the catheter with an inner tube disposed within an outer tube and defining a first flow path interiorly of the inner tube and second flow path between the inner tube and the outer tube. This inserted catheter also includes a plurality of hollow fibers disposed in fluid communication with the first and second flow paths. The method further includes steps for creating a flow of heat exchange fluid through the first and second flow paths, and moving the inner tube relative to the outer tube to change the profile of the hollow fibers.

In a further aspect of the invention, a heat exchange catheter includes an elongate shaft with first portions defining an inlet lumen and second portions defining an outlet lumen. A first manifold is disposed in fluid communication with the inlet lumen and a second manifold disposed in fluid communication with the outlet lumen. A plurality of hollow fibers are disposed between the manifolds in fluid communication with the inlet and outlet lumens. The catheter is adapted to receive a heat exchange fluid and to direct the heat exchange fluid through the hollow fibers to exchange heat through the hollow fibers.

In still a further aspect of the invention, a catheter is adapted to exchange heat with the body fluid flowing in a first direction through a body conduit. The catheter includes a shaft having an input lumen and an output lumen. A plurality of hollow fibers define a heat exchange region and collectively define an output surface of the heat exchange region. The input lumen of the shaft is coupled to the hollow fibers at a first location while the output lumen of the shaft is coupled to the hollow fibers at a second location disposed in the first direction from the first location.

Another aspect of the invention includes a method for exchanging heat with a body fluid in a body conduit. In this case, a catheter is provided with a plurality of hollow heat exchange fibers extending in fluid communication with an inlet lumen and an outlet lumen of the catheter. The heat exchange fibers collectively define a first cavity in heat transfer relationship with a body fluid in a body conduit.

In an additional aspect of the invention, an operative area of the catheter is sized and configured for disposition in a vessel containing blood. The operative area is adapted to perform a predetermined function, and the blood in the vessel has a tendency to form clots. In this aspect of the invention, the catheter is provided with a snare disposed relative to the operative area and being operable from a proximal end of the catheter to move from a low-profile state facilitating insertion of the catheter, to a high-profile state facilitating the capture of blood clots.

In still a further aspect of the invention, a heat exchange catheter is adapted for cooling the blood of a patient. The catheter includes a heat exchange region with a plurality of fibers each having a hollow configuration. A heat exchange fluid is disposed in the hollow fibers to cool the fibers and a coating is disposed on the outer surface of the fibers to inhibit formation of blood clots.

These and other features and advantages of the invention will be better understood with a description of the preferred embodiments of the invention and reference to the associated drawings.

Description of the Drawings

Fig. 1 is side elevation view of a patient lying in a prone position with a heat exchange catheter of the present invention appropriately inserted to facilitate hypothermic treatment of the patient's brain;

Fig. 2 is an enlarged side elevation view showing the vasculature associated with the patient's head and brain;

Fig. 3 is a perspective view partially in section of a heat exchange region of the catheter;

Fig. 4 is an enlarged axial cross section view of a plurality of balloons disposed in the heat exchange region of the catheter;

Fig. 5 is a radial cross section view of the catheter taken along lines 5-5 of Fig. 4;

Fig. 6 is a radial cross section view similar to Fig. 5 of a further embodiment of the catheter;

Fig. 7 is a perspective view of a further embodiment of the catheter wherein multiple balloons are provided with a longitudinal configuration;

Fig. 8 is a radial cross section view taken along lines 8-8 of Fig. 7;

Fig. 9 is an axial cross section view taken along lines 9-9 of Fig. 7;

Fig. 10 is a perspective view of the catheter illustrated in Figure 3 further illustrating structures which can facilitate mixing and heat exchange;

Fig. 10A is a perspective view of an embodiment of the catheter having a distal end with a pigtail configuration;

Fig. 10B is a perspective view of the catheter illustrated in Fig. 10A with the distal end straightened by a stylet 174 to facilitate insertion of the catheter;

Fig. 11 is a schematic view of an embodiment including a heat pipe;

Fig. 12 is a schematic view, partially in section, of a heat pipe adapted for use in the embodiment of Fig. 11;

Fig. 13 is a top plan view of carotid artery branch illustrating one method of operation associated with the catheter;

Fig. 14 is a top plan view similar to Fig. 13 and showing a further method of operation with the catheter;

Fig. 15 is a top plan view of the carotid branch similar to Fig. 13 and showing a further method of operating a heat exchange catheter;

Fig. 16 is a radial cross section of the catheter taken along lines 16-16 of Fig. 15;

Fig. 17 is an axial cross section view of a further embodiment of the invention including hollow fibers in the heat exchange region;

Fig. 18 is a side elevation view similar to Figure 17 and illustrating the hollow fibers in a compacted configuration; and

Fig. 19 is an axial cross section view of the catheter of Figure 17 operatively disposed and configured to permit the hollow fibers to float and undulate within a blood stream.

Fig. 20 is a side elevation view partially in section and illustrating a further embodiment of the catheter of the present invention;

Fig. 21 is a radial cross-section view taken along the lines 21-21 of Figure 20;

Fig. 22 is an axial cross-section view of the proximal end of the catheter illustrated in Figure 20;

Fig. 23 is an axial cross-section view of the distal end of a further embodiment illustrating the heat exchange region in a low-profile state;

Fig. 24 is an axial cross-section view similar to Figure 23 and illustrating the heat exchange region in a high-profile state;

Figs. 25-27 illustrate a preferred method for manufacturing the heat exchange region of a hollow fiber embodiment of the cavity;

Fig. 25 is a top plan view of a mat formed of the heat exchange fibers;

Fig. 26 is a perspective view illustrating formation of the mat around the distal ends of the concentric tubes;

Fig. 27 is a side elevation view illustrating attachment of the mat assembly to an outer tube of the catheter;

Fig. 28 is a top-plan view of a patient illustrating portions of the blood circulatory system;

Fig. 29-33 illustrate a method for introducing the catheter of the present invention;

Fig. 29 is a side elevation view illustrating a introducing sheath in a first position removed from the heat exchange region;

Fig. 30 is a side elevation view illustrating the sheath in a second position over the heat exchange region of the catheter;

Fig. 31 is a side elevation view illustrating the catheter and sheath being inserted into an introducer;

Fig. 32 is a side elevation view illustrating the catheter further inserted with the sheath maintained in the introducer;

Fig. 33 is a side elevation view illustrating removal of the sheath to the first position;

Fig. 34 is a perspective view of a further embodiment of the catheter including a distal clot filter in a low-profile state;

Fig. 35 is a perspective view illustrating the catheter of Figure 34 with the clot filter in a high-profile state;

Fig. 36 is a perspective view of a catheter with a clot filter having free ends and automatically deployable to a high-profile state; and

Fig. 37 is a side elevation view of the catheter of Figure 36 with a sheath maintaining the clot filter in a low-profile state.

**Description of the Preferred Embodiments and
Best Mode of the Invention**

A heat exchange catheter is illustrated in Figure 1 and designated generally by the reference numeral 10. The catheter 10 is operatively disposed with respect to a body 12 of a patient having a groin 14, a head 16, and a brain 18. More specifically, the catheter 10 can be inserted percutaneously through a puncture or surgical cut down at the groin 14, and into the femoral artery 21. Following this initial introduction, the catheter 10 can be moved through the femoral artery 21 and the aortic arch 23, into the common carotid artery 25 best illustrated in Figure 2. This common carotid artery 25 divides at a carotid branch 27 into an external carotid artery 30, which primarily supplies blood 31 to the face of the patient, and an internal carotid artery 32, which primarily supplies blood to the brain 18 of the patient.

In the concept of this invention, the brain 18 is merely representative of a portion of the body 12 of the patient, and the arteries 21, 25, 30 and 32 are merely representative of conduits which supply a body fluid, such as blood, to a selected portion of the body 12, such as the brain 18. By cooling the body fluid, such as blood 31, in the body conduit, such as the artery 32, the specific body portion, such as the brain 18, can be selectively cooled without significantly affecting the temperature of the remaining portions of the body 12.

Selective hypothermic treatment of the brain 18 is initially of particular interest as it captures the advantages of hypothermia during operative procedures associated with the brain 18 without also capturing the disadvantages of hypothermia with respect to other areas of the body 12. Thus, a surgeon operating to treat an aneurysm in the brain 18, for example, can initially cool the brain 18 in order to facilitate that procedure. This selective hypothermia will be particularly appreciated in those surgical procedures which are primarily directed to the brain 18. Procedures such as stroke, trauma, and other brain related injuries will also benefit up to and during from this selective hypothermia treatment.

A preferred embodiment of the catheter 10 of the present invention is illustrated in Figure 3 and 4. From this perspective view, it can be seen that the catheter 10 includes a shaft

40 having an axis 41 which extends between a proximal end 43 and a distal end 45. When operatively disposed, a heat exchange region 47 at the distal end 45 is operatively disposed within the body 12, and a hub 50 at the proximal end 43 is disposed outside of the body 12. Within the shaft 40, a plurality of lumens 52 and 54 extend in fluid communication with the hub 50 and the heat exchange region 47.

A preferred embodiment of the heat exchange region 47 is illustrated in greater detail in Figure 4 where three balloons 56, 58 and 61 are individually, separately and axially disposed along the shaft 40. It will be appreciated that although the illustrated embodiment includes three balloons, a single balloon or double balloon embodiment may offer further advantages in a particular procedure. All of the balloons 56, 58 and 61 are illustrated to have a significantly larger diameter than the shaft 40. This may not be the case in other embodiments. More specifically, it may be desirable to maximize the dimension of the shaft 40 in order to facilitate flow of the heat exchange fluid. This will also minimize the volume of fluid in the balloon and promote a more rapid heat exchange. In one such embodiment, the diameter of the shaft 40 is in a range between 50 and 90 percent of the diameter of the balloons 56, 58 and 61.

Each of the balloons 56, 58 and 61 can be formed from a piece of sheet material 62, 64 and 66 which is bound or otherwise fixed to the shaft 40 to form a cavity 63, 65 and 67, respectively. An inlet hole 70 provides fluid communication between the lumen 54 and the cavity 63 of the balloon 56. Similar inlet holes 72 and 74 are provided for the balloons 58 and 61. In a like manner, an outlet hole 76 can be formed in the wall of the shaft 40 to provide fluid communication between the lumen 52 and the cavity 63 of the balloon 56. Similar outlet holes 78 and 81 are provided for the balloons 58 and 61, respectively. With this structure, it can be seen that the lumen 54 functions primarily as an inlet lumen for a heat exchange fluid which is illustrated generally as a series of arrows designated by the reference numeral 85.

Initially, the heat exchange fluid 85 is introduced through the hub 50 (Figure 3) and into the inlet lumen 54. From the lumen 54, the heat exchange fluid 85 passes through the

After the heat exchange fluid 85 is remotely cooled, it is circulated through the balloon cavities 63, 65 and 67 to provide a cold temperature fluid on the inner surface of the sheet materials 62, 64 and 66 which form the walls of the balloons 56, 58 and 61, respectively. With a body fluid, such as blood 31, flowing exteriorly of the balloons 56, 68 and 61, heat transfer occurs across the sheet materials 62, 64 and 66, respectively.

It can be appreciated that this circulation of the heat exchange fluid 85 can be formed with any structure of the shaft 40 which provides two lumens, such as the lumens 52 and 54, each of which can have access to the balloon cavities, such as the cavities 63, 65 and 67. In one embodiment of the shaft 40 illustrated in Figure 5, a septum 90 is provided which separates the cylindrical shaft 40 into two equally sized lumens 52 and 54. In the embodiment of Figure 6, the cylindrical shaft 40 is provided with a cylindrical septum 92 which provides the lumen 54 with a circular cross section and the lumen 52 with a moon-shaped cross section. In such an embodiment, the lumen 54 must be defined off-axis from the shaft 40 in order to have access to the balloon cavities, such as the cavity 63.

One of the advantages of a multiple balloon embodiment of the catheter 10 is that the flow and temperature of the heat exchange fluid 85 can be more easily controlled along the entire length of the heat exchange region 47. Realizing that the heat exchange fluid 85 will be coolest prior to entering into a heat exchange with the blood 31, and warmest after that heat exchange, one can advantageously control not only the velocity and volume of flow, but also the direction of flow within each discrete balloons 56, 58 and 61. Another advantage of a multiple balloon design is the ability of the catheter to bend and flex when placed in a curved vasculature. Single balloon designs would be rigid, stiff and inflexible by comparison.

In order to facilitate the maximum heat exchange between the fluid 85 and the blood, it is desirable to provide a balanced flow of the heat exchange fluid 85 along the entire length

of the heat exchange region 47. In the embodiment illustrated in Figure 4, efficient heat transfer is facilitated by countercurrent flow where the heat exchange fluid 85 is directed to flow counter to the flow of the blood 31. To that end, the inlet holes 70, 72 and 74 are positioned distally of the outlet holes 76, 78 and 81, respectively. As the blood 31 flows distally along the outer surface of the catheter 10, this relative position of the inlet holes and outlet holes causes the heat exchange fluid to flow in the opposite direction, proximally in each of the balloons 56, 58 and 61.

The amount of flow within each of the balloons 56, 58 and 61 can also be controlled by the size of the inlet holes 70, 72, 74 and outlet holes 76, 78 and 81. In a preferred embodiment, this flow control is provided solely by the inlet holes 70, 72 and 74; the outlet holes 76, 78 and 81 are sized larger than their respective inlet holes so that they offer little resistance to flow. In this embodiment, the inlet holes 70, 72 and 74 are sized to be progressively smaller from the distal end 45 to the proximal end 43. Thus the hole 70 is larger than the hole 72 which is larger than the hole 74. As a result, the resistance to the flow of heat exchange fluid 85 in the most distal balloon 56 is less than that in the most proximal balloon 61. This ensures that the coolest heat exchange fluid 85 is shared equally among all of the balloons 56, 58 and 61 regardless of their position along the shaft 40. In an embodiment wherein the flow is controlled by the outlet holes 76, 78 and 81, these holes can also be provided with a relatively reduced size from the distal end 45 to the proximal end 43. With any of these structures, a more balanced flow of the heat exchange fluid can be achieved in order to facilitate the highest degree of heat exchange along the entire heat exchange region 47. Alternatively, the flow of heat exchange fluid can also be balanced by providing the holes 76, 78 and 81 with non-circular configurations. For example, these holes may be formed as longitudinal slits extending axially of the catheter.

A further embodiment of the invention is illustrated in Figure 7 wherein a single sheet of material 101 is used to form separate and distinct individual balloons, two of which are designated by the reference numerals 103 and 105. As opposed to the radial balloons 56, 58

and 61 of the previous embodiment, the balloons 103 and 105 extend axially along the surface of the shaft 40. For example, the balloons 103 and 105 form individual balloon cavities 107 and 110, respectively, which extend from a distal end 112 to a proximal end 114.

This embodiment of the catheter containing the axial balloons 103 and 105 may include a shaft 40 with a slightly different configuration. As best illustrated in Figure 9, the shaft 40 may include an outer tube 121 having an outer surface to which the sheet material 101 is attached and within which is disposed a distal sealing plug 123. An inner tube 125, which can be disposed coaxially with the outer tube 121, has an inner lumen 127 and defines with the outer tube 121 an outer lumen 130. A pair of inlet holes 132 and 134 provide flow fluid communication between the inner lumen 127 and the balloon cavities 107 and 110, respectively. Similarly, a pair of outlet holes 136 and 138 provide fluid communication between the balloon cavities 107 and 110 and the outer lumen 130, respectively. An inner plug 141 disposed between the inner tube 125 and outer tube 121 to seal the outer lumen 130 between the inlet holes 132, 134 and outlet holes 136, 138. For the reasons previously noted, a preferred embodiment has inlet holes 132, 134 which are disposed distally of and sized smaller than the outlet holes 136, 138, respectively. This orientation will provide countercurrent flow in a catheter 10 which is inserted downstream into an artery such as the carotid artery 25.

Embodiments which are intended to maximize heat transfer will take advantage of the fact that heat exchange is enhanced when either, or both, the body fluid or the heat exchange fluid is provided with well mixed flow. Mixing can be enhanced by providing irregular surfaces next to which either of these fluids flow. For example, with reference to Figure 4, it will be noted that a spring 150 can be disposed around the shaft 40 inside each of the balloons, such as the balloon 61. In this embodiment, the spring 150 upsets the laminar flow of the heat exchange fluid 85 thereby producing the desired mixing of this fluid. Other structures can be positioned within the cavities formed by the balloons 56, 58 and 61.

Mixing can also be enhanced within the body fluid which flows along the outer surface of the catheter 10. In this case, the multiple radial balloon embodiment illustrated in Figure 4 is of advantage as each of the balloons 56, 58 and 61 represents a peak and defines with the adjacent balloon a valley along which the blood 31 flows. This series of peaks and valleys also upsets the laminar flow of the body fluid. Mixing of the body fluid can also be enhanced by providing other structures along the outer surface of the sheet material 62, 64 and 66 which form the balloons as well as any exposed areas of the shaft 40 in the heat exchange region 47. By way of example, a multiplicity of granules 145 can be adhered to the outer surface of the radial balloons 56, 58 and 61 or the axial balloons 103 and 105 as illustrated in Figure 9. Ridges can also be provided along these surfaces.

With some body fluids, it may be desirable to inhibit turbulent flow and facilitate laminar flow. This may be true for example in the case of blood where undesirable hemolysis may occur in response to increased turbulence. Such an embodiment might be particularly desirable for use with radial balloons where an outer balloon 152 would promote laminar flow by reducing the height differential between the peaks and valleys defined by the balloons 56, 58 and 61. This outer balloon 152 is best illustrated in Figure 10. To further promote laminar flow, the outer surface of any structure in the heat exchange region 47 can be provided with a coating 154, such as a hydrophilic or a hydrophobic coating to modify the boundary layer. Thus the outer surface of the shaft 40 as well as the outer surface of any of the balloons 56, 58, 61, 103, 105 and 152 can be provided with the coating 154. The coating 154 may also include other ingredients providing the catheter 10 with additional advantageous properties. For example, the coating 154 may include an antithrombogenic ingredient such as heparin or aspirin. Such a coating 154 would not only inhibit platelet deposition but also the formation of blood clots.

As previously noted, the characteristics of the heat exchange fluid 85 may also be of importance in a particular heat exchange environment. Although the heat exchange fluid 85 may include various liquids, it is believed that gases may provide the greatest temperature

differential with the body fluid. Particularly if this fluid includes blood, gases that are inert or otherwise compatible with the vascular system will be appreciated. Although several inert gases might fulfill these requirements, carbon dioxide is used for the heat exchange fluid 85 in a preferred embodiment of the invention.

A further embodiment of the catheter 10 is contemplated for maximizing the surface area available for heat exchange. As illustrated in Figures 10A and 10B, the catheter 10 can be formed with a distal end 45 of the shaft 40 disposed in the natural configuration of a spiral or pigtail 172. The relatively large diameter of the pigtail 172 facilitates heat exchange, but tends to deter from a low profile desire for insertion. Under these circumstances, it may be advantageous to insert the catheter 10 over a stylet or guidewire 174 in order to straighten the pigtail 172 as illustrated in Figure 10B.

Hyperthermia and hypothermia for selective regions of the body can also be achieved by placing in the body conduit, such as the carotid artery 25, a heat pipe 161 best illustrated in the schematic view of Figure 11. In this embodiment, the heat pipe 161 includes a distal end 163 and proximal end 165. The distal end 163 is adapted to be placed within the body conduit, such as the carotid artery 25. The proximal end 165 of the heat pipe 161 is adapted to be connected to an external heat sink or cooler, such as a thermoelectric cooler 167 or water jacket 168. A wick structure 170 is provided in the heat pipe 161 to facilitate a flow of heat exchange fluid from the cooler 167 to the distal end 163.

In a process involving the heat pipe 161, illustrated in Figure 12, the heat exchange fluid is moved from the proximal end 165 of the heat pipe 161 either by gravity or by capillary action of the wick structure 170 to the distal end 163. At the distal end 163 of the heat pipe 161, heat is transferred from the body fluid, such as blood, to the heat exchange fluid in its liquid state. This heat exchange liquid absorbs a heat of vaporization as it passes into a vapor state in the heat pipe 161. The heat exchange fluid in its vapor state creates a pressure gradient between the ends 163 and 165 of the heat pipe 161. This pressure gradient causes the vapor to flow to the cooler 165 where it is condensed giving up its latent heat of

vaporization. The heat exchange fluid in its liquid state then passes back through the heat pipe 161 through the wick structure 170 or by gravity. The passive heat exchange system provided by the heat pipe 161 is vacuum-tight and can be operated with a minimum amount of the heat exchange fluid.

Although the heat exchange catheter 10 will be advantageous in the hyperthermic or hypothermic treatment of any portion of the body 12, it is believed that it will be particularly appreciated in those procedures which can benefit from the hypothermic treatment of the brain 18, such as the treatment of ischemic stroke and/or head trauma. As previously noted in comments directed to Figure 1, the catheter 10 can be inserted into the femoral artery in the groin 14 and directed through the aortic arch 23 into the common carotid artery 25. As illustrated in Figure 13, the catheter 10 can then be moved into the region of the arterial branch 27 where it will encounter the external carotid artery 30 and the internal carotid artery 32. Since the external carotid artery 30 is directed primarily to the facial regions, it does not supply a significant amount of blood to the brain 18. In contrast, the internal carotid artery 32 is almost solely responsible for feeding the capillary bed of the brain 18. Based on these considerations, hypothermic treatment of the brain 18 is best addressed by cooling the blood in the internal carotid artery 32 without wasting any of the cooling properties on the external carotid artery 30. In a method associated with one embodiment of the invention, the most distal of the balloons, such as the balloon 56 in Figure 13 is preferably positioned within the internal carotid artery 32. The more proximal balloons 58 and 61 can be disposed along the common carotid artery 25. This embodiment of the catheter 10 and its associated method will achieve a higher degree of heat transfer within the internal artery 32 than the external artery 30.

In another embodiment of the catheter 10 best illustrated in Figure 14, an occlusion balloon 175 is provided distally of the heat exchange region 47. In this embodiment, the occlusion balloon 175 will preferably be inflatable through a separate lumen in the shaft 40. As the catheter 10, approaches the carotid branch 27, the occlusion balloon 81 is directed into

the external carotid artery 30 and inflated in order to at least partially occlude that artery. The remaining proximal balloons 56, 58 and 61 in the heat exchange region 47 are left within the common carotid artery 25 to promote heat exchange with the blood flowing to the branch 27. With the external artery 30 at least partially occluded, heat transfer occurs primarily with the blood flowing into the internal carotid artery 32.

A further embodiment of the invention is illustrated in Figure 15 operatively disposed in the common carotid artery 25 and internal carotid artery 32. In this case, the catheter 10 includes a balloon 181 which is attached to the distal end of the shaft 40 and provided with a spiral configuration. More specifically, the balloon 181 may be formed from several individual balloons, as with the embodiment of Figure 7, for as individual flutes 183 on the single balloon 181. In either case, the separate balloons (such as the balloons 103, 105 of Figure 7) or the flutes 183 are oriented in a spiral configuration around the axis 41 of the catheter 10. The shaft 40 can be provided with any of the configurations previously discussed such as the eccentric configuration of Figure 6.

By providing the balloon 181 with a spiral configuration, heat exchange is enhanced by at least two of the factors previously discussed. Notably, the surface area of contact is increased between the blood 31 flowing externally of the balloon 181 and the heat exchange fluid flowing internally of the balloon 181. The spiral configuration also enhances the mixing properties of both the blood 31 and the heat exchange fluid 85.

As noted, the heat exchange fluid 85 may be cooled to a sub-zero temperature. In order to thermally protect the internal lining of the artery 32 from direct contact with the sub-zero coolant, it may be desirable to provide the tips of the flutes 183 with a thicker wall 185, as shown in Figure 16.. This thicker wall 185 might be advantageous in any of the balloon configurations previously discussed, but would appear to be most advantageous in the embodiments of Figure 7 and 15 where the contact with the artery 32 tends to be more localized by the longitudinal balloons 103, 105 (Figure 7) on the spiral flutes 183 (Figure 15).

Still a further embodiment of the invention is illustrated in Figure 17. In this embodiment, the shaft 40 includes an inner tube 190 disposed within an outer tube 192. These tubes 190, 192 may be concentric and longitudinally movable relative to each other. The tubes 190, 192 terminate respectively in manifolds 194, 196. Between these manifolds 194, 196, a multiplicity of hollow fibers 198 can be disposed at the distal end 45 to define the heat exchange region 47 of the catheter 10. The hollow fibers 198 each include an internal lumen which provides fluid communication between the manifolds 194 and 196. In operation, the heat exchange fluid 85 flows distally along the inner tube 190 into the distal manifold 194. From this manifold 194, the heat exchange fluid 85 flows into the internal lumens of the hollow fibers 198 proximally to the proximal manifold 196. The warmer heat exchange fluid 85 flows proximally from the manifold 196 between the inner tube 190 and outer tube 192.

Preferably, the hollow fibers 198 have a wall thickness that is thin enough to allow maximum heat transfer, yet strong enough to withstand the pressure requirements of the heat exchange fluid 85. The hollow fibers 198 are further adapted to achieve ideal heat transfer by the maximization of both surface area and coolant flow. The smaller the diameter of the fibers 198, the more fibers can be fit into the catheter 10 with a corresponding increase in surface area. As the diameter of the fibers 198 is decreased, however, the resistance to fluid flow increases thus lowering the coolant flow rate. The effect of the inflow and outflow lumens must also be considered in determining the fluid resistance. Ideally, the wall thickness of the hollow fibers 198 is in a range between .00025 inches and .003 inches. In a preferred embodiment the wall thickness is in a range between .00075 inches and .002 inches, and ideally .00125 inches. The outer diameter of the hollow fibers 198 will typically be between .008 inches and .035 inches. In a preferred embodiment the outer diameter is in a range between .010 inches and .018 inches, and ideally .015 inches.

It will be noted that the heat exchange fluid 85 flowing in the inner tube 190 is insulated in several respects from the blood stream outside the catheter 10. This flow channel in the inner tube 190 is insulated not only by the wall of the outer tube 192, but also by the

coolant returning in the flow channel associated with the outer tube 192. The heat exchange fluid 85 in the inner tube is further insulated by the thickness of the inner tube wall.

In the heat exchange region 47, the wall thicknesses associated with the inner tube 190 and the outer tube 192 is preferably reduced in order to provide additional volume for the hollow fibers 198. With a reduced wall thickness, the inner tube 190 also contributes to the heat exchange occurring in the region 47.

The hollow fibers 198 offer several advantages to this embodiment of the catheter 10. Notably, they provide a very high surface area between the blood 31 and the heat exchange fluid 85. This greatly enhances the heat exchange characteristics of this embodiment. Countercurrent flow can also be maintained further facilitating the heat exchange capabilities of this catheter.

The hollow fibers 198 can be spiraled as illustrated in Figure 18 by twisting the inner tube 190 with respect to the outer tube 192. This characteristic can be used to provide a shorter and lower profile heat exchange region 47 in order to facilitate introduction of the catheter 10. A lower profile may also be obtained by separating the manifolds 194 and 196 a distance substantially equal to the length of the fibers 198. This will tend to hold the fibers in a straight, parallel relationship and thereby facilitate introduction of the catheter 10. The spiraled configuration of the hollow fibers 198 can be maintained during heat exchange in order to further increase the heat exchange area per unit length of the catheter 10. Alternatively, the fibers 198 can be positioned to loosely float and undulate between the manifolds 194 and 196 as illustrated in Figure 19. This characteristic of the fibers 198 will not only provide the increased heat exchange area desired, but also promote mixing within the blood 31.

The fibers 198 will typically be formed of common materials such as polyolefin nylon and polyurethane. The fibers can be coated with a clot-inhibiting material such as heparin. Other materials advantageous for inhibiting the formation of blood clots might include those which form polymer surfaces with 16 or 18 carbon alkyl chains. These materials attract

albumin and thereby inhibit clot formation. In a further embodiment, the fibers 198 can be provided with micropores which permit the leaching of such clot inhibiting pharmaceuticals as heparinized saline which could also serve as the heat exchange fluid 85.

The embodiment of Figure 20 also takes advantage of the significant heat exchange characteristics associated with the hollow fibers 198. In this embodiment, the manifold 194 at the distal end 45 of the catheter 10 includes a potting seal 201 with a distal surface 203. The fibers 198 are held in the potting seal 201 with the lumens of the fibers 198 exposed at the surface 203. The distal end of the inner tube 190 is also held in the potting seal 201 with its lumen exposed at the distal surface 203. In this embodiment, the manifold 194 includes a cap 205 which may have a hemisphere configuration. This cap extends over the distal surface 203 of the potting seal 201 and provides fluid communication between the lumen of the inner tube 190 and the lumens of the hollow fibers 198. This cap 205 may also be constructed of materials and wall thicknesses that insulate the blood vessels from potential contact with a cold catheter tip.

Figure 21 illustrates in a cross-sectional view a first flow channel 204 which extends along the lumen of the inner tube 190 and a second flow channel 206 which extends along the lumen of the outer tube 192 outwardly of the inner tube 190. As the heat exchange fluid 85 is introduced into the first flow channel 204, its direction is reversed in cap 205 so that the flow of the fluid 85 in the hollow fibers is counter to the flow of the body fluid, such as blood, in the body conduit, such as the artery 32. After moving through the fibers 198, the heat exchange fluid 85 passes along the second flow channel 206 between the inner tube 190 and outer tube 192, and exits the catheter 10 at the proximal end 43.

The embodiment of Figure 20 also includes a Y-connector 207 disposed at the proximal end 43 of the catheter 10. This connector 207 is shown in greater detail in the enlarged view of Figure 22. In this view it can be seen that the connector 207 includes a body 210 with screw threads 212 at its distal end and screw threads 214 at its proximal end. At

the distal end of the body 210, a screw cap 216 mates with the screw threads 212 to engage an annular flange 218 at the proximal end of the outer tube 192. In this manner, the Y-connector 207 forms a seal with the proximal end of the outer tube 192 and provides fluid communication between the second flow channel 206 and a lumen 221 of the Y-connector 207. A side port 223 communicates with this lumen 221 and provides an exit port for the secondary flow channel 206.

In order to prevent leakage from the lumen 221 at the proximal end 43 of the Y-connector 207, a releasable seal 225 can be formed at the proximal end of the body 210. In the illustrated embodiment, the releasable seal 225 includes a cap 227 which is threaded to register with the threads 214 of the body 210. This cap 227 extends around the proximal end of the body 210 and compresses an elastomeric washer 230 against the body 210 and the outer surface of the inner tube 190. By tightening the cap 227, the washer 230 is compressed to seal the lumen 221. This compression also functions to inhibit, but not necessarily prevent, axial movement between the outer tube 192 and inner tube 190. The releasability of the seal 225 can be appreciated in order to facilitate this relative movement between the tubes 190 and 192 for the reasons previously discussed. This form of a releasable seal 225 is commonly referred to as a Tuohy-Borst seal.

The relative movement between the inner and outer tubes 190 and 192, respectively, will be appreciated in order to provide the tubes 190 and 192 with a first position wherein the fibers 198 have a low profile configuration as illustrated in Figure 23. The relative movement will also be appreciated in order to provide the tubes 190 and 192 with a second position wherein the hollow fibers 198 form an increased profile as illustrated in Figure 24. It can be appreciated that this profile will facilitate heat exchange by providing an increased spacing of the individual hollow fibers in the body fluid.

Another feature associated with these two positions is illustrated in Figure 23 where the inner tube 190 is expanded in thickness at its distal end in order to form a ramp or taper 232. In this embodiment, the taper 232 is annular and extends radially outward with

progressive distal positions along the tube 190. As the inner tube 190 is drawn proximally relative to the outer tube 192, the taper 232 is brought into sealing engagement with the proximal end of the hollow fibers 198 and potting seal 201. This effectively seals the distal end of the outer tube 192 against the outer surface of inner tube 190, and prohibits any loss of the heat exchange fluid 85 between the inner and outer tubes 190 and 192 at the distal end 45.

This loss of the heat exchange fluid 85 can also be addressed with a seal tube 234 which can be positioned between the inner and outer tubes 190, 192 and inwardly of the hollow fibers 198. In this embodiment, a distal end 236 of the seal tube 234 is generally coextensive with the distal end of the outer tube 192. The seal tube 234 is preferably provided with an inner diameter greater than the outer diameter of the inner tube 190. As a result, the inner tube 190 is free to move relative to the outer tube 192 to achieve the advantages previously discussed. However, when the inner tube 190 is drawn sufficiently proximal of the outer tube 192, the taper 232 will contact the distal end 236 of the seal tube 234. This effectively forms the seal between the inner and outer tubes 190 and 192, respectively at the distal end of the outer tube 192. With the taper 232 wedged against the seal tube 234, the fibers 198 are maintained in their operative free-floating configuration as illustrated in Figure 24.

Alternatively, a non-tapered inner tube 190, can be mated with a closely fitted seal tube 234. With very small and controlled differences between the outside diameter of the inner tube 190 and the inside diameter of the seal tube 234, for example 0.0005 to 0.003 inches, an effective seal can be constructed without the taper 232. This embodiment relies on the length of the seal tube 234, the surface tension of the coolant fluid 85, and the small capillary gap to create a resistance greater than the pressure of the coolant fluid during operation. This design does not require the inner tube to be moved a fixed distance relative to the outer tube and does not require a constant tension between the inner and outer tubes to effect a seal.

The seal tube 234 is preferably constructed of polyimide which allows for a precision and constant inner diameter. In addition, polyimide is available in very thin wall thicknesses so that the seal tube 234 will not occupy a significant portion of the annular space which is more appropriately dedicated to the fibers 198.

A method for manufacturing the hollow fiber embodiments of the catheter 10 is illustrated in Figures 25-27. In Figure 25, a planar mat 241 of the hollow fibers 198 is formed with a generally planar configuration. In this mat 241, the fibers 198 are oriented in a generally parallel configuration with angled potting seals 201 and 243 formed at opposite ends of the fibers 198. This fiber mat 241 can be rolled onto the outer surfaces of the inner tube 190 and seal tube 234 as illustrated in Figure 26. In this step, the potting seal 201 is formed around the distal end of the inner tube 190 while the potting seal 243 is formed around the distal end of the seal tube 234.

By initially forming the fibers 198 into the mat 241, a generally uniform thickness of the mat 241 can be maintained. Rolling the mat 241 onto the tubes 190 and 234 maintains this uniform thickness and also facilitates orientation of the fibers 198 onto the cylindrical tubes 190 and 234. This technique also forms an inwardly spiraling helical bond joint profile that aids in directing the blood flow in order to inhibit clot formation by preventing stagnant blood flow areas at the bond joint. With the potting seals 201 and 243 suitably bonded to the tubes 190 and 234, respectively, the cap 205 can be mounted over the distal end of the fibers 198 as previously discussed. At the proximal end of the fibers 198, the seal tube 234 can be mounted in the distal end of the outer tube 192 as illustrated in Figure 27.

The seal tube 234 offers some interesting possibilities for the infusion of fluids at the distal end 45 of the catheter 10. Of course, it is always possible to provide an additional lumen within the shaft of the catheter 10. In such an embodiment, the fluid to be infused could be injected into the additional lumen at the proximal end 43 to exit the catheter at the distal end 45. Alternatively, the fluid to be infused might be included in the heat exchange fluid 85. The tolerance between the seal tube 234 and the outer diameter of the inner tube

190 could then be controlled to provide a calibrated leak of the heat exchange fluid 85 at the distal end 45 of the catheter 10. Micro holes might also be drilled into the outer tube 192 or inner tube 190 to provide for a controlled leakage of the infusion fluid.

Each of the foregoing embodiments of the heat exchange catheter 10 is adapted for use in cooling the entire human body, or perhaps only a portion of the total body. Methods of operation will vary widely depending on the focus of a particular procedure. By way of example, it will be noted with reference to Figure 28 that the catheter 10 is particularly adapted for cooling blood in a procedure which may involve as many as three of the catheters 10. In Figure 28, a human body 245 is illustrated along with a portion of the blood circulatory system including a pair of femoral veins 247, 250 and a subclavian vein 252. These veins 247, 250 and 252 all extend into the vena cava 254 of the body 245. In this procedure, separate catheters, such as the heat exchange catheter 10, can be introduced into each of the femoral veins 247, 250 and the subclavian vein 252 with their respective heat exchange regions disposed in the vena cava 254. Alternatively, and preferably, only two such catheters would be introduced from two of the three veins 247, 250 and 252.

A systemic version of the catheter might have a diameter in a range of between 9 and 15 French, and a length of approximately 20 to 80 centimeters long. It is contemplated that this design could conceivably cool the body in several hours. The use of two such catheters inserted into the vena cava 254 as mentioned above could be expected to reduce the time required to cool the body by a factor of 2. It will be appreciated that similar catheters and methods can be used to lower the temperature of blood in the native carotid or in the vertebral circulatory system. The amount of blood heat lost is directly proportional to the temperature differential, the blood velocity and the blood-to-catheter surface area.

Particularly in an operative setting wherein the heat exchange catheter 10 is to be inserted into a blood vessel, a further design feature best illustrated in Figures 29-33 will be of particular interest. In these views, an introducer 256 is positioned for percutaneous insertion into a blood vessel such as the femoral vein 250. A sleeve 258 is provided on the

catheter 10 and slidable along the outer tube 192 between two positions. The first position is illustrated in Figure 29 wherein the sleeve 258 is disposed in a spaced relationship with the heat exchange region 47. The second position of the sleeve 258 is illustrated in Figure 30 where the sleeve 258 covers the heat exchange region 47. In this position the balloons or fibers associated with the region 47 are compressed to a low profile state greatly facilitating introduction of the catheter 10 into the introducer 256. In addition, the covered heat exchange region 47 is stiffened for easier introduction into the introducer 256. The fibers and/or balloons are also protected from the interior surface of the introducer 256. Optionally, a stiffening mandril may be inserted down one or more of the tubes 190, 192 to facilitate introduction of the catheter 10 into the introducer 256.

After this initial insertion, the sleeve 258 remains within the introducer 256 while the remainder of the heat exchange region 47 is moved distally into the conduit as illustrated in Figure 31. At this point, the sleeve 258 can be removed from the introducer 256 by sliding it proximally to its first position as illustrated in Figure 33.

This method of introduction is facilitated by providing the sleeve 258 with a generally cylindrical configuration. The diameter of the cylindrical sheath should be less than the inside diameter of the introducer 256. However, at the proximal end of the sheath 258, an annular flange 261 or other enlargement can be provided to ensure that the sheath 258 does not pass beyond the introducer 256.

Another feature associated with the present invention relates to a blood clot basket or snare 263, best illustrated in Figures 34 and 35. This snare 263 is preferably positioned downstream of the heat exchange region 47 associated with the catheter 10. It being appreciated that any structure disposed in a blood vessel may tend to generate blood clots, it is the purpose of the snare 263 to capture any such clots. The snare 263 of the preferred embodiment includes a plurality of wires 265 which extend along a shaft 267 with their opposing ends fixed in the manifold 194 and a distal cap 270. The wires 265 in a preferred embodiment are formed of stainless steel or a nickel titanium alloy.

In the illustrated embodiment, the shaft 267 extends to the proximal end 43 of the catheter 10 either through the lumen of the inner tube 190 or alternatively through a second, separate lumen in the inner tube 190. In the former case, a seal would be required at the distal end of the manifold 194 to prevent any leakage of heat exchange fluid 85 around the shaft 267.

In either case, the shaft 267 is free to move relative to the concentric tubes 190 and 192. When the shaft 267 is moved relatively distally, the snare wires 265 are provided with a generally low profile. When the shaft 267 is moved relatively proximally, the wires 265 deploy to provide the snare with an enlarged high-profile configuration as illustrated in Figure 35.

In a further embodiment of the snare 263, the wires 265 are connected to the manifold 194 and extend to distal ends which are unattached or free. The wires 265 in this embodiment, best illustrated in Figure 36, are bent to a deployed enlarged configuration. With such an embodiment, insertion is facilitated by providing a delivery sheath which is movable to maintain the wires 265 in a low-profile state. Once the catheter 10 is in place, the sheath 262 can be removed thereby permitting the wires 265 to automatically expand to their enlarged high-profile state.

With respect to the forgoing disclosure as a whole, it will be apparent that many variations from these preferred embodiments will now be apparent to those skilled in the art. For example, with respect to the balloon embodiments previously discussed, it will be appreciated that the advantages of this invention can be derived with only a single balloon. On the other hand, there seem to be several advantages associated with multiple balloon embodiments. Notably, a more even and balanced transfer of heat exchange can be achieved with multiple balloons. In addition, there appears to be better mixing with respect to both the blood 31 as well as the heat exchange fluid 85. Multiple balloons also provide an increased surface area relative to single balloon embodiments. Furthermore, the overall flexibility of the catheter 10 is enhanced with multiple balloons separated by interruptions which provide

natural flex points for the catheter. When the balloons experience the high perfusion pressure, they become more stiff. The reduced diameter interruptions provide for increased flexibility at these joints.

Additional flexibility can be derived by providing the shaft 40 with variable stiffness. This variability can be produced by different materials forming the shaft 40 along its length or alternatively, tapering or otherwise varying the diameter of the shaft 40. For example, the shaft 40 can be progressively tapered from its proximal end 43 to its distal end 45 in order to provide a softer and more flexible heat exchange region 47.

In any of the foregoing embodiments of the catheter 10, the inner tube 190 can be provided with a central lumen facilitating introduction over a guidewire and providing a capability for the infusion of fluids through the catheter 10.

With the intent of maximizing heat transfer with the body fluid in a conduit feeding a specific region of the body, any of the factors previously noted can be addressed to provide structural modifications to the foregoing embodiments. Of course changes in the material or size of any of the structural elements described can be varied to achieve various heat exchange properties. Realizing the many changes which might be contemplated, one is cautioned not to limit this concept only to the specific embodiments illustrated and disclosed, but rather to determine the scope of the invention with reference to the following claims.

Claims

1. A catheter having an elongate configuration with a proximal end and a distal end, the catheter comprising:
 - an outer tube having an elongate configuration and a first lumen;
 - an inner tube disposed in the first lumen of the outer tube and having a second lumen extending between the proximal end and the distal end of the catheter;
 - portions of the inner tube defining a first fluid flow path extending along the second lumen between the proximal end and the distal end of the catheter;
 - portions of the outer tube and the inner tube defining a second flow path extending between the first tube and the second tube; and
 - a plurality of hollow fibers providing fluid communication between the first fluid flow path and the second fluid flow path.
2. The catheter recited in Claim 1, wherein:
 - each of the hollow fibers has a proximal end and a distal end;
 - the distal end of each of the hollow fibers has a fixed relationship with the distal end of the inner tube; and
 - the proximal end of each of the hollow fibers has a fixed relationship with the distal end of the outer tube.
3. The catheter recited in Claim 2, wherein the inner tube has properties for moving relative to the outer tube to vary the configuration of the hollow fibers extending between the inner tube and the outer tube.
4. The catheter recited in Claim 3, wherein:

portions of the inner tube define a taper, the inner tube being axially movable to bring the portions of the inner tube into sealing proximity with the hollow fibers.

5. The catheter recited in Claim 1 further comprising:
a cap disposed over the distal ends of the inner tube and the hollow fibers.
6. The catheter recited in Claim 1, further comprising:
a seal tube disposed inwardly of the proximal end of the hollow fibers and forming a seal with the distal end of the outer tube and the proximal ends of the hollow fibers.
7. The catheter recited in Claim 6, wherein the seal tube extends proximally of the proximal end of the hollow fibers.
8. The catheter recited in Claim 4, further comprising:
a seal tube disposed inwardly of the proximal end of the hollow fibers and forming a seal with the distal end of the outer tube and the proximal end of the hollow fibers; and
the portions of the inner tube which define the taper are axially movable relative to the outer tube in the sealing engagement with the hollow fibers.
9. The catheter recited in Claim 1, wherein the hollow fibers are adapted to receive a heat exchange fluid from the first flow path and to release the heat exchange fluid into the second flow path.
10. The catheter recited in Claim 3, further comprising:
a seal tube disposed between the hollow fibers and the inner tube and having an inner diameter greater than the outer diameter of the inner tube, but sufficiently close to the outer

diameter of the inner tube to form a liquid seal between the seal tube and the inner tube by capillary action.

11. The catheter recited in Claim 10, wherein:
portions of the inner tube define a taper, the inner tube being axially movable to bring the portions of the inner tube into sealing proximity with the distal end of the seal tube.
12. The catheter recited in Claim 5, further comprising:
a coating of insulation covering the cap at the distal end of the catheter.
13. The catheter recited in Claim 1, wherein the hollow fibers are adapted to receive a heat exchange fluid from the second flow path and to release the heat exchange fluid into the first flow path.
14. A method for making a heat exchange catheter, comprising the steps of:
providing a first tube having a first lumen extending between a proximal end and a distal end;
inserting a second tube into the lumen of the first tube, the second tube having a second lumen;
connecting a plurality of hollow fibers in fluid communication with a first flow path extending along the second lumen of the second tube, and a second flow path extending along the first lumen of the first tube outwardly to the second tube; and
insuring that the second tube is at least one of axially and rotationally movable relative to the first tube to vary the configuration of the hollow fibers in order to facilitate heat exchange with the heat exchange catheter.
15. The method recited in Claim 14, wherein the insuring step includes the steps of :

moving the second tube distally relative to the first tube to change the hollow fibers to a low profile state; and

moving the second tube proximally relative to the first tube to change the hollow fibers to a high profile state.

16. The method recited in Claim 14, wherein the connecting step further comprises the steps of:

forming the hollow fibers in a stack having a generally planar configuration;
wrapping the hollow fibers stack around the second tube; and
inserting the hollow fibers into the distal end of the first tube.

17. The method recited in Claim 14, further comprising the steps of:

fixing to the proximal end of the first tube a Y-connector having fluid communication with the second flow path.

18. The method recited in Claim 17, further comprising the steps of:

attaching a locking device to the Y-connector, the locking device being operable between a first position permitting movement of the second tube relative to the first tube, and a second position inhibiting movement of the second tube relative to the first tube.

19. The method recited in Claim 16, further comprising the step of:

potting the hollow fiber stack to form an end seal tapered radially inwardly to inhibit formation of stagnant flow regions around the fibers of the stack.

20. A method for operating a heat exchange catheter within a body conduit containing a body fluid, the method comprising the steps of:

inserting into the body conduit the heat exchange catheter with an inner tube disposed within an outer tube to define a first flow path interiorly of the inner tube and a second flow path between the inner tube and the outer tube, and a plurality of hollow fibers disposed in fluid communication between the first flow path and the second flow path; and

creating a flow of heat exchange fluid by introducing the heat exchange fluid into one of the first flow path and the second flow path.

21. The method recited in Claim 20, wherein the inserting step includes the step of pushing the inner tube distally relative to the outer tube to provide the hollow fibers with a lower profile.

22. The method recited in Claim 20, wherein the creating step further comprises the step of reciprocating the inner tube relative to the outer tube to create a continuous movement of the hollow fibers in order to facilitate heat exchange between the hollow fibers and the body fluid.

23. The method recited in Claim 20, further comprising the steps of:
removing the catheter from the body conduit, and prior to the removing step, pushing the inner tube distally relative to the outer tube to provide the hollow fibers with a lower profile.

24. The method recited in Claim 20, wherein prior to the inserting step the method comprises the steps of:

providing a sheath outwardly of the outer tube and movable between a first position spaced from the hollow fibers and a second position proximate to the hollow fibers; and

moving the sheath to the second position and over the hollow fibers to maintain the fibers in a low profile state.

25. The method recited in Claim 24, further comprising the steps of after the inserting step, moving the sheath from the second position to the first position.
26. The method recited in Claim 25, further comprising the step of after the first moving step, moving the sheath from the first position to the second position and withdrawing the catheter.
27. The method recited in Claim 20, wherein the heat exchange fluid is a cooling fluid.
28. A heat exchange catheter, including:
an elongate shaft extending along an axis between a proximal end a distal end;
first portions of the shaft defining an inlet lumen extending between the proximal end and the distal end of the shaft;
second portions of the shaft defining an outlet lumen;
a first manifold in fluid communication with the inlet lumen at the distal end of the shaft;
a second manifold in fluid communication with the outlet lumen of the shaft;
a plurality of hollow fibers disposed to extend between the first manifold and the second manifold in fluid communication with the inlet lumen and the outlet lumen; and
the catheter being adapted to receive a heat exchange fluid at the proximal end of the inlet lumen, and to direct the heat exchange fluid through the hollow fibers to exchange heat through the hollow fibers.
29. The heat exchange catheter recited in Claim 28, wherein the first manifold is disposed distally of the second manifold.

30. The heat exchange catheter recited in Claim 29, wherein the outlet lumen is disposed outwardly of the inlet lumen.
31. The heat exchange catheter recited in Claim 28, wherein the shaft comprises:
an inner tube defining the input lumen; and
an output tube concentric with the input tube and defining with the input tube the output lumen.
32. The heat exchange catheter recited in Claim 31, wherein:
the first tube has a fixed relationship with the first manifold;
the second tube has a fixed relationship with the second manifold; and
the first tube is axially movable relative to the second tube to vary the configuration of the hollow fibers.
33. The heat exchange catheter recited in Claim 32, wherein the first tube is movable axially of the second tube to separate the first manifold and the second manifold, and to place the hollow fibers in a generally straight, parallel relationship.
34. The heat exchange catheter recited in Claim 28, wherein the heat exchange fluid is a liquid.
35. A catheter adapted to exchange heat with a body fluid flowing through a body conduit, the catheter comprising:
a shaft having an axis extending between a proximal end and a distal end, the shaft having an input lumen and an output lumen;
a plurality of hollow fibers defining a heat exchange region of the shaft and collectively defining an outer surface of the heat exchange region;

36. The catheter recited in Claim 35, wherein the body fluid flows in a first direction through a body conduit and the heat exchange fluid flows through the hollow fibers in a second direction opposite to the first direction.

38. The catheter recited in Claim 36, further comprising:
a clot snare disposed in the first direction from the heat exchange region.

40. A method for exchanging heat with a body fluid in a body conduit, comprising the steps of:

providing the catheter with a first cavity in heat transfer relationship with a body fluid in the body conduit;

introducing a heat exchange fluid into the inlet lumen and into the first cavity;
exchanging heat between the heat exchange fluid and the body fluid in the body conduit;

removing the heat exchange fluid from the first cavity through the outlet lumen; and
during the providing step, providing the catheter with a plurality of hollow heat exchange fibers each extending in fluid communication with the inlet lumen and the outlet lumen, the heat exchange fibers collectively defining the first cavity in heat transfer relationship with the body fluid in the body conduit.

41. The method recited in Claim 40, wherein the second introducing step includes the step of introducing the heat exchange fluid into the hollow fibers.

42. A catheter having a elongate configuration with a proximal end and distal end, comprising:

an operative area of the catheter sized and configured for disposition in a vessel containing blood flowing in a particular direction, the operative area being adapted to perform a predetermined function and the blood in the vessel having a tendency to form blood clots; and

a snare disposed in the particular direction from the operative area and being operable from the proximal end of the catheter to move from a low-profile state facilitating insertion of the catheter into the vessel, and a high-profile state facilitating capture of any blood clots.

43. The catheter recited in Claim 42, wherein the snare is disposed distally of the operative area of the catheter.

44. The catheter recited in Claim 42, wherein the operative area includes a heat exchange region of the catheter.

45. The catheter recited in Claim 44, wherein the heat exchange region is a heat receiving region of the catheter.
46. The catheter recited in Claim 42, wherein the snare includes:
a plurality of elongate filaments each having a first end attached to the operative region of the catheter and a second end.
47. The catheter recited in Claim 46, wherein the second ends of the filaments are unattached.
48. The catheter recited in Claim 46, wherein:
the catheter further comprises a cap disposed at the distal end of the catheter;
the second ends of the filaments are attached to the cap; and
movement of the cap relative to the operative region of the catheter changes the profile of the snare.
49. The catheter recited in Claim 48, further comprising:
a shaft attached to the cap and extending to the proximal end of the catheter; whereby
the profile of the snare is changed by movement of the shaft relative to the operative region of the catheter.
50. The catheter recited in Claim 42, wherein the filaments are formed of wire.
51. The catheter recited in Claim 50, wherein the wire filaments include a nickel titanium alloy.

52. A heat exchange catheter having an elongate configuration and extending between a proximal end and a distal end, the catheter being adapted for cooling the blood of a patient, comprising:

- a heat exchange region of the catheter;
- a plurality of fibers included in the heat exchange region, with each of the fibers having a hollow configuration and being adapted to receive a heat exchange fluid; and
- a coating disposed on the outer surface of the fibers to inhibit the formation of blood clots on the cooled fibers.

53. The heat exchange catheter recited in Claim 52, further comprising a chemical included in the coating and having characteristics for inhibiting the formation of the blood clots.

54. The heat exchange catheter recited in Claim 53, wherein the chemical includes heparin.

55. The heat exchange catheter recited in Claim 53, wherein:
each of the fibers includes a multiplicity of micro pores extending between the hollow interior of the fibers and the outer surface of the fibers; and
the chemical is included in the heat exchange fluid and leached with the heat exchange fluid through the micro pores to coat the outer surface of the fibers.

56. The method recited in Claim 20 further comprising the step of:
moving the inner tube relative to the outer tube to change the profile of the hollow fibers.

57. The method recited in Claim 20, wherein the creating step includes the step of:

providing a heat exchange fluid in the form of a liquid.

58. The method recited in Claim 20, wherein the creating step includes the step of:
providing a heat exchange fluid in the form a gas.
59. The method recited in Claim 20, wherein the creating step includes the step of:
heating the heat exchange fluid prior to introducing the heat exchange fluid into the
catheter.
60. The method recited in Claim 20, wherein the creating step includes the step of:
cooling the heat exchange fluid prior to introducing the fluid into the catheter.
61. The heat exchange catheter recited in Claim 28, wherein the heat exchange fluid is a
gas.
62. The heat exchange catheter recited in Claim 28, wherein the heat exchange fluid is a
cooling fluid.
63. The heat exchange catheter recited in Claim 28, wherein the heat exchange fluid is a
heating fluid.

ABSTRACT

**INDWELLING HEAT EXCHANGE CATHETER
AND METHOD OF USING SAME**

A catheter is adapted to exchange heat with a body fluid, such as blood, flowing in a body conduit, such as a blood vessel. The catheter includes a shaft with a heat exchange region disposed at its distal end. This region may include hollow fibers which are adapted to receive a remotely cooled heat exchange fluid preferably flowing in a direction counter to that of the body fluid. The hollow fibers enhance the surface area of contact, as well as the mixing of both the heat exchange fluid and the body fluid. The catheter can be positioned to produce hypothermia in a selective area of the body or alternatively positioned to systemically cool the entire body system.

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August 13, 1998

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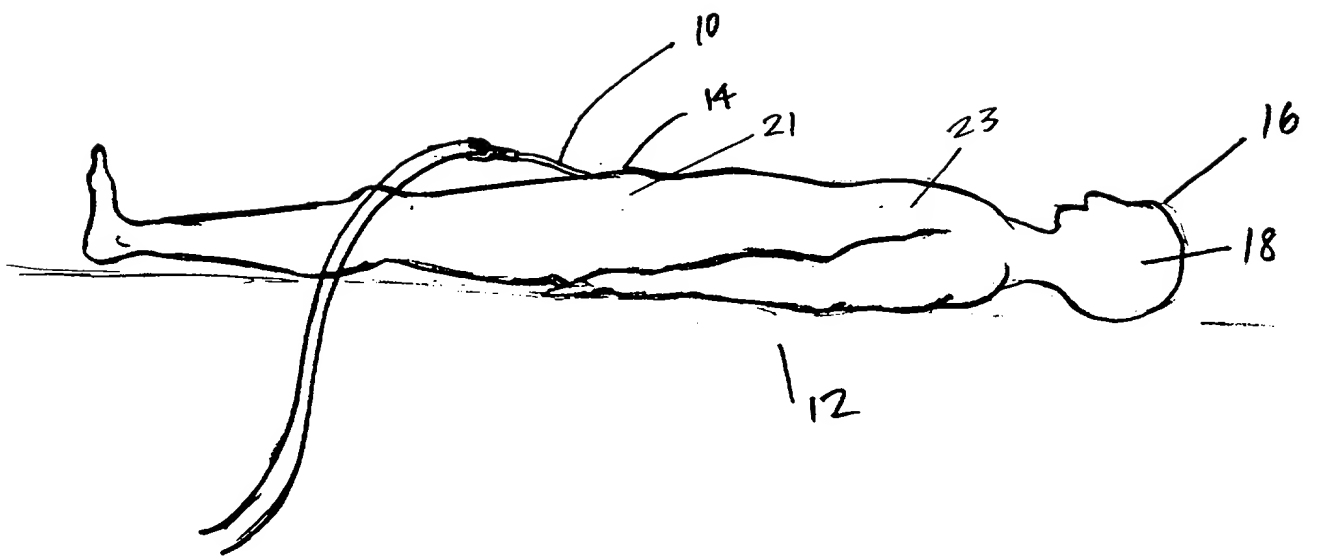


FIG. 1

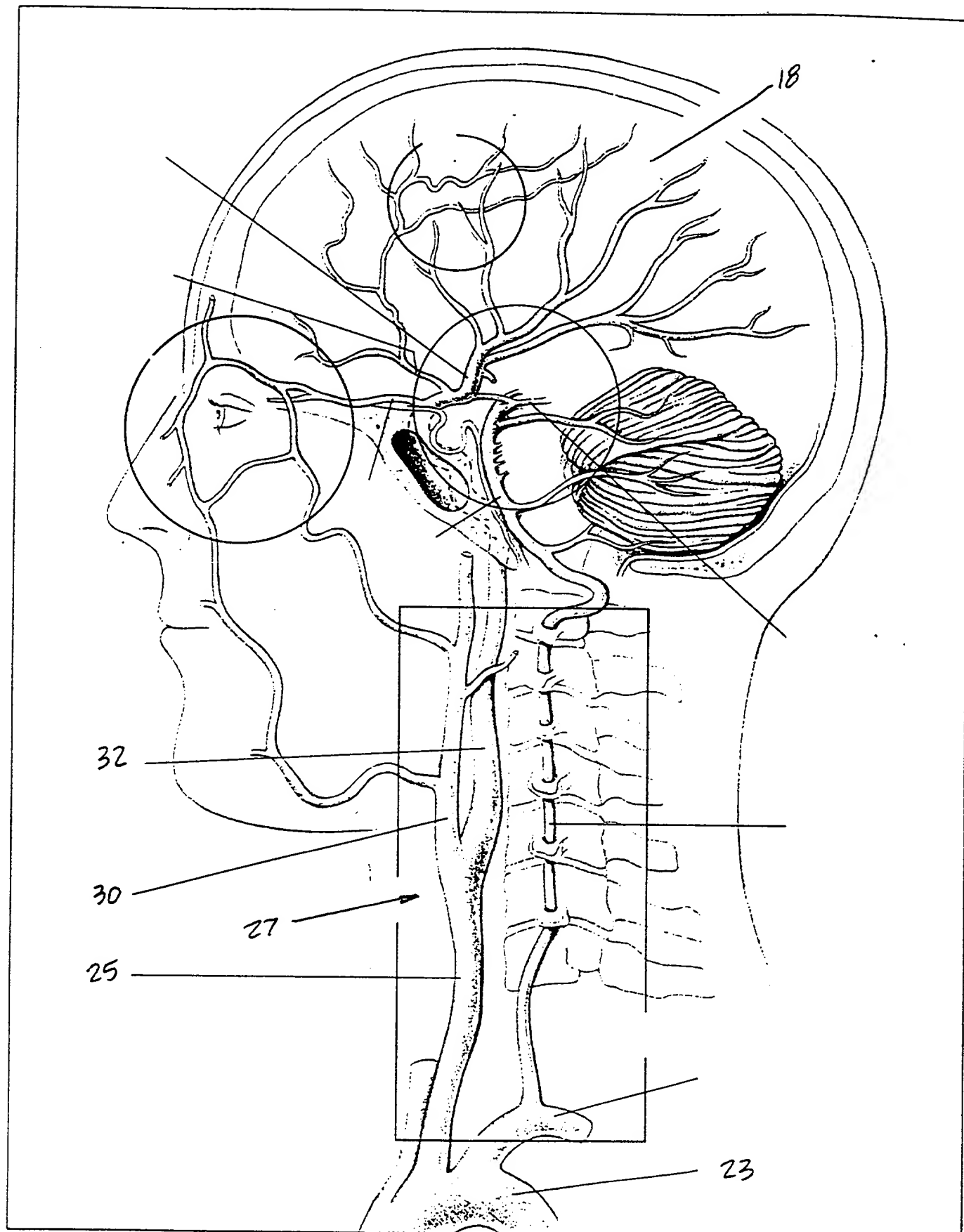


FIG. 2

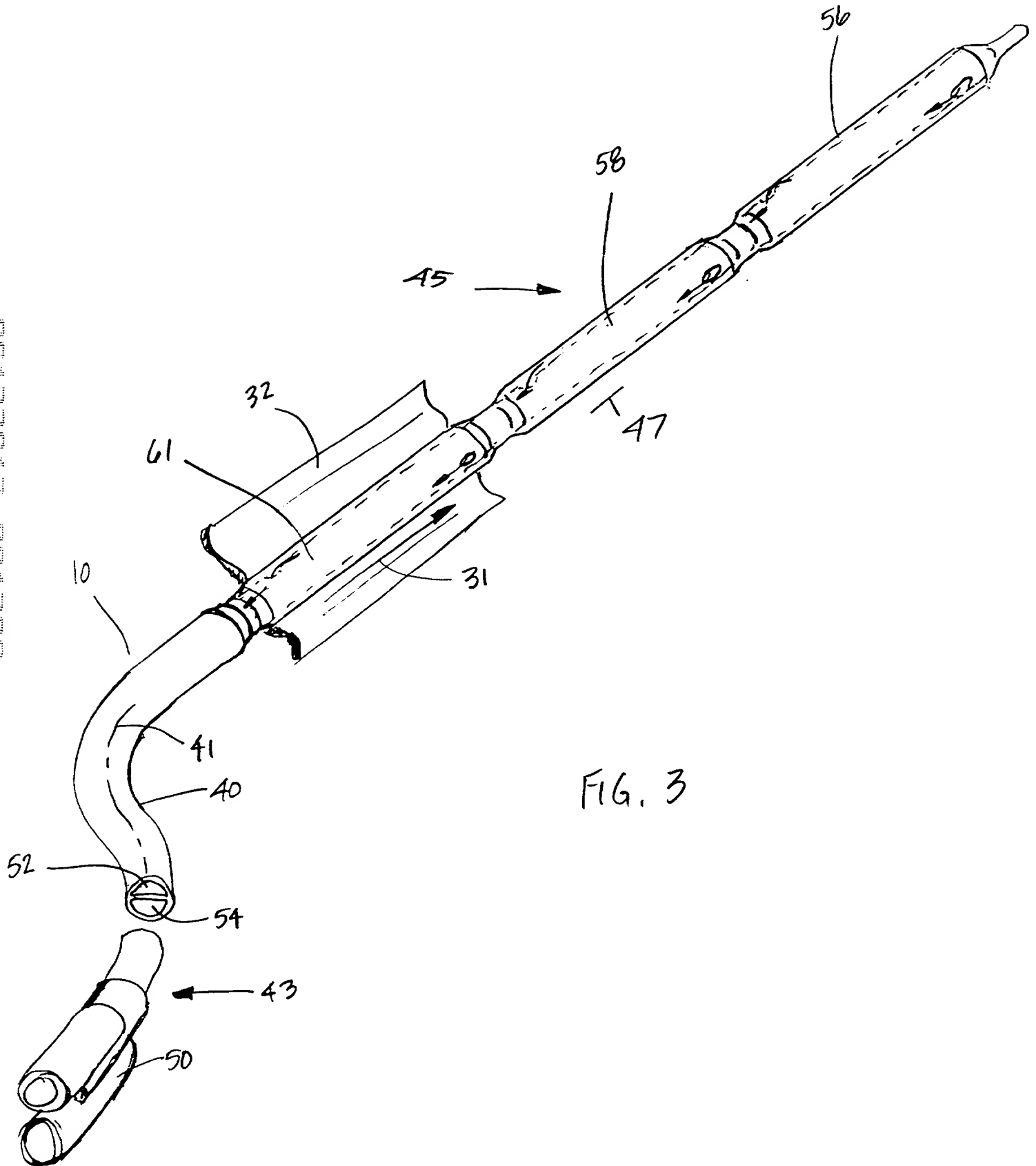
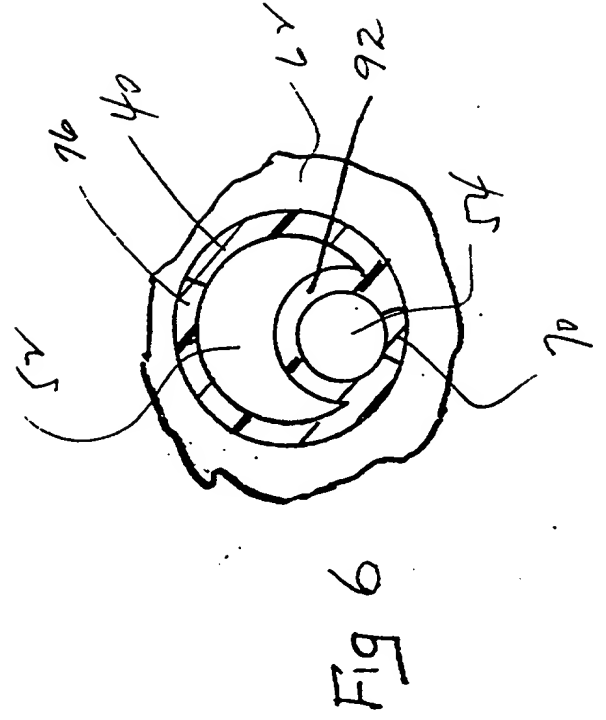
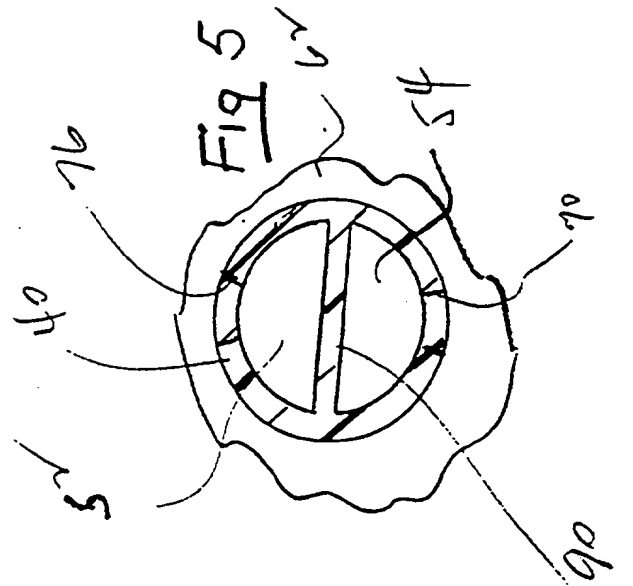
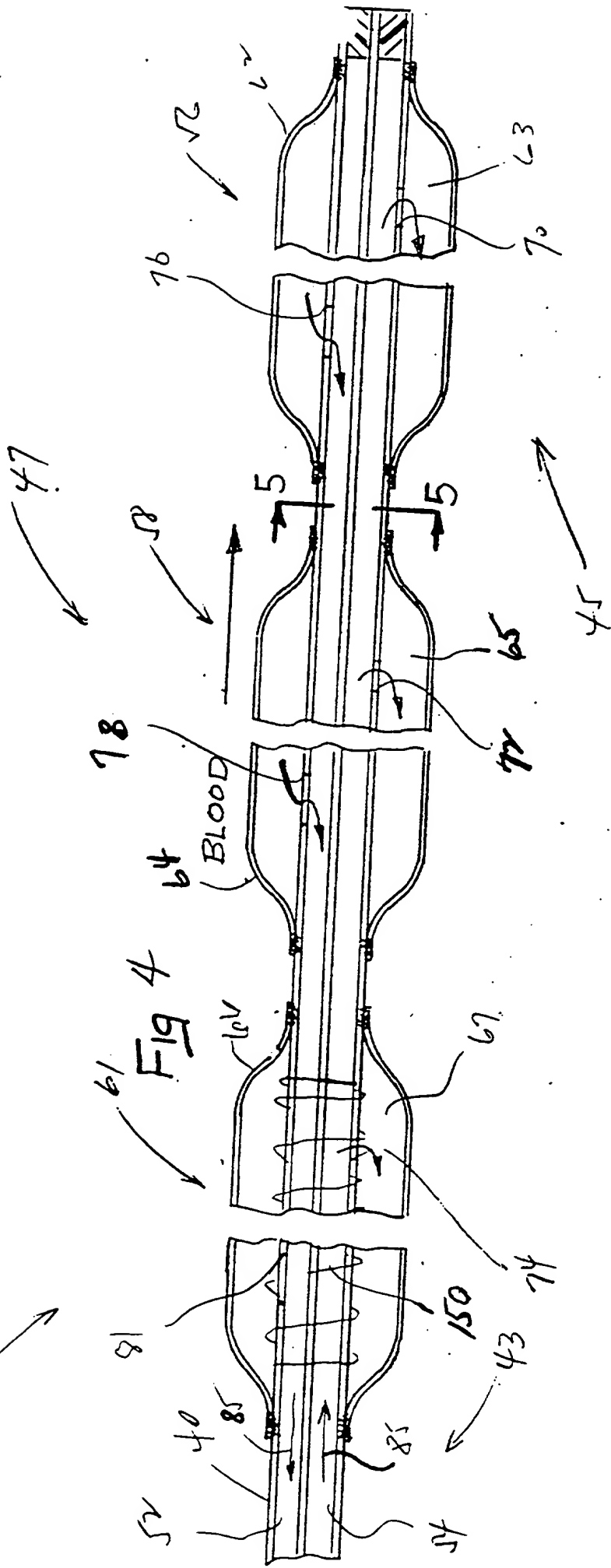
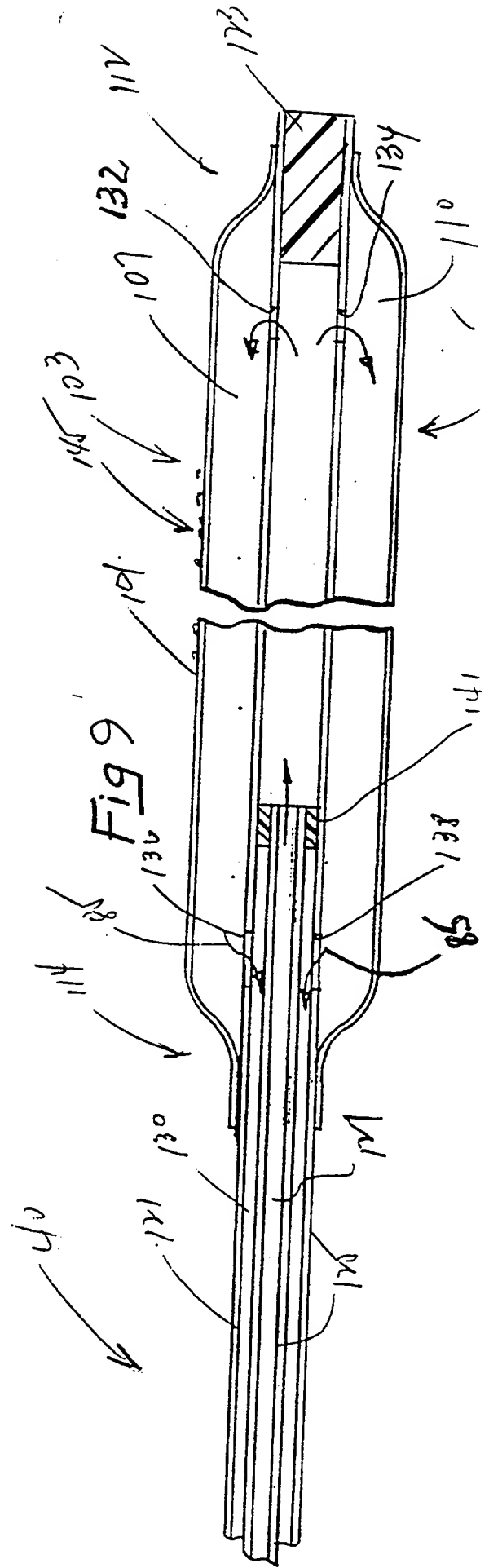
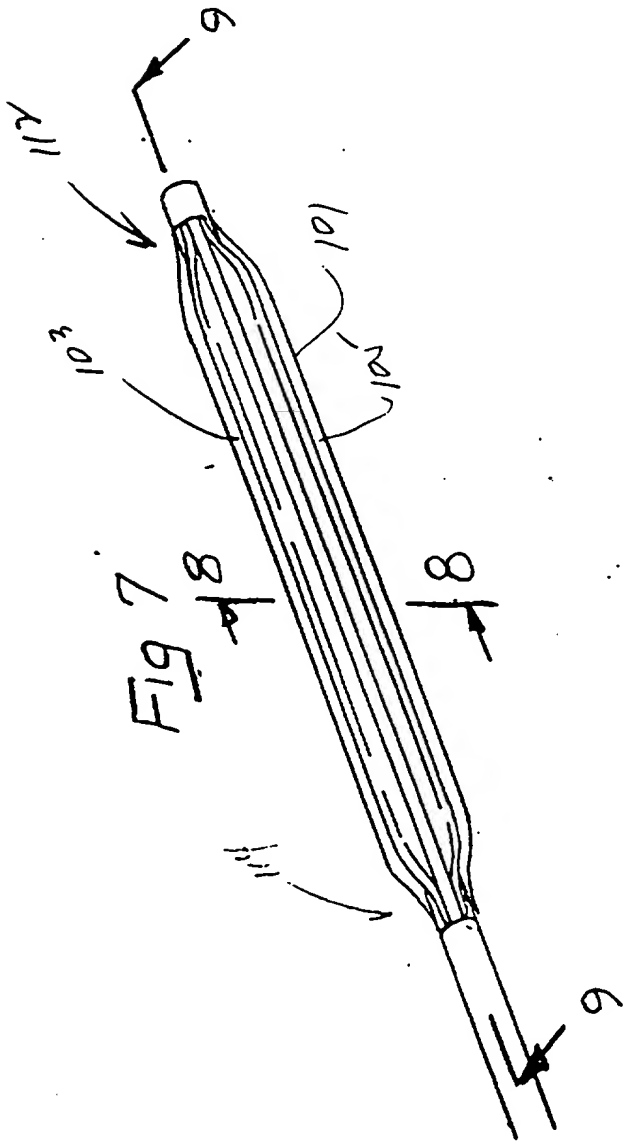
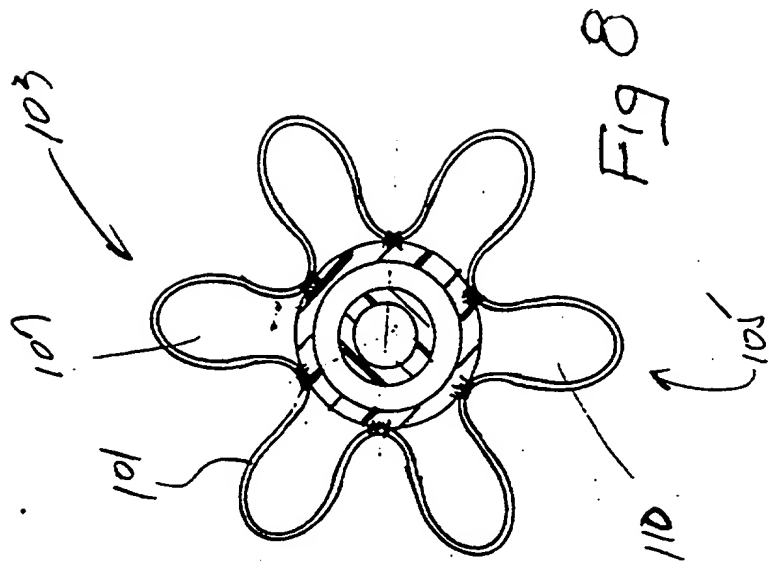
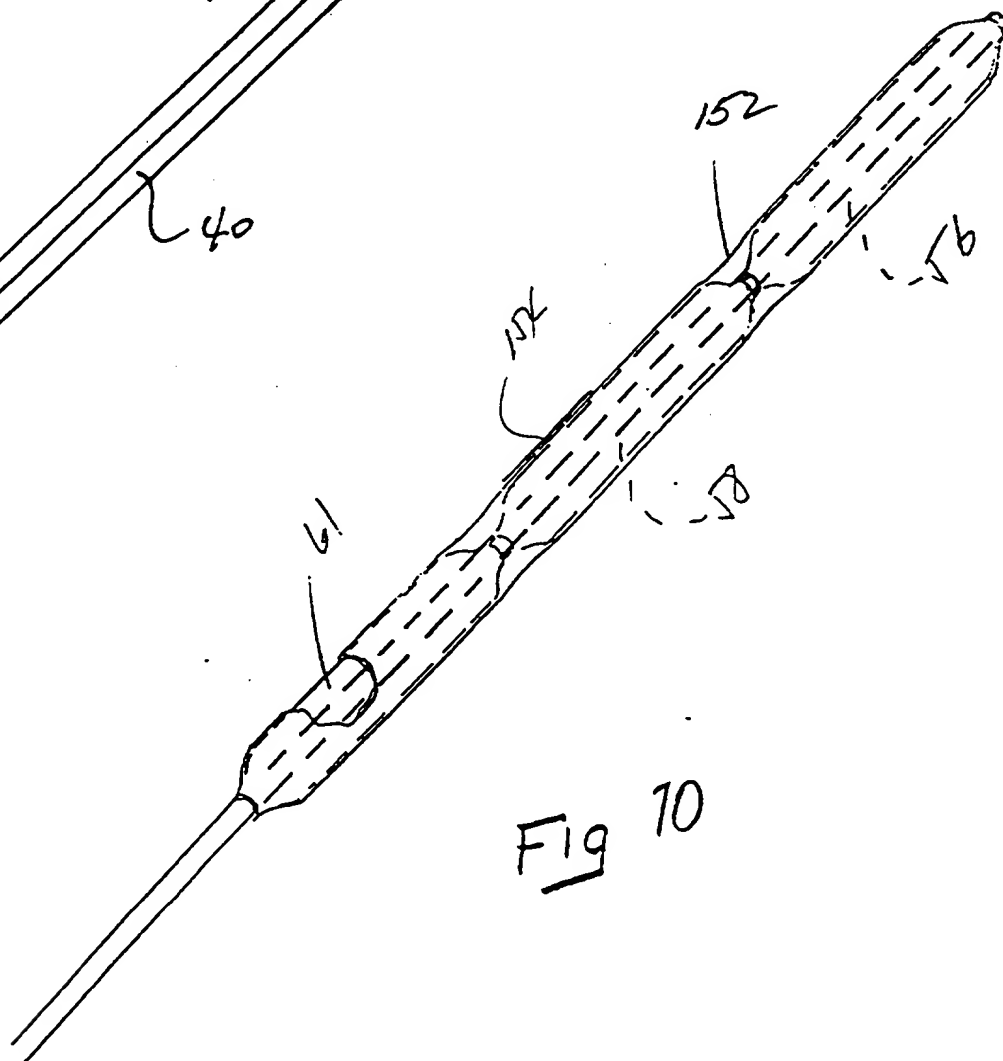
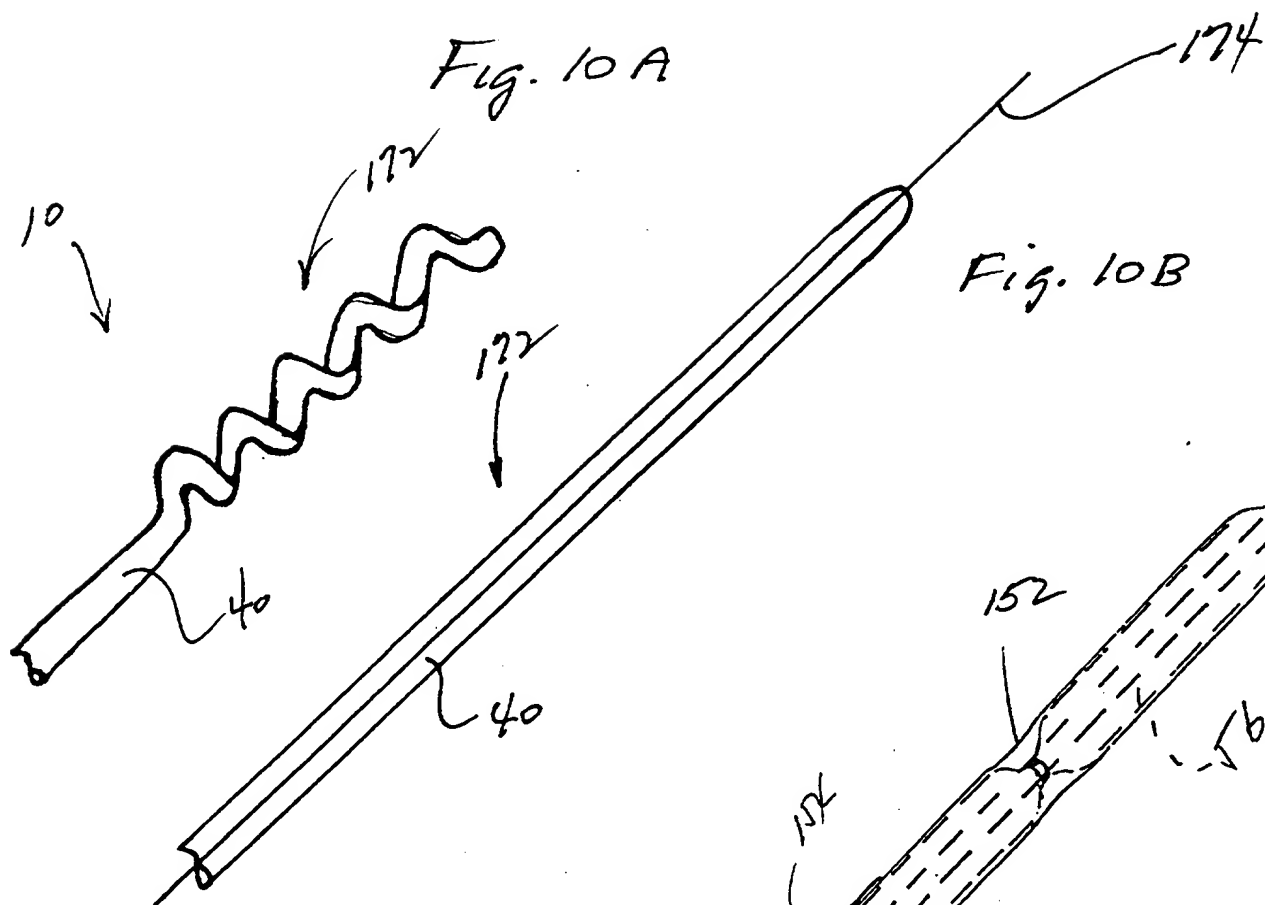


FIG. 3







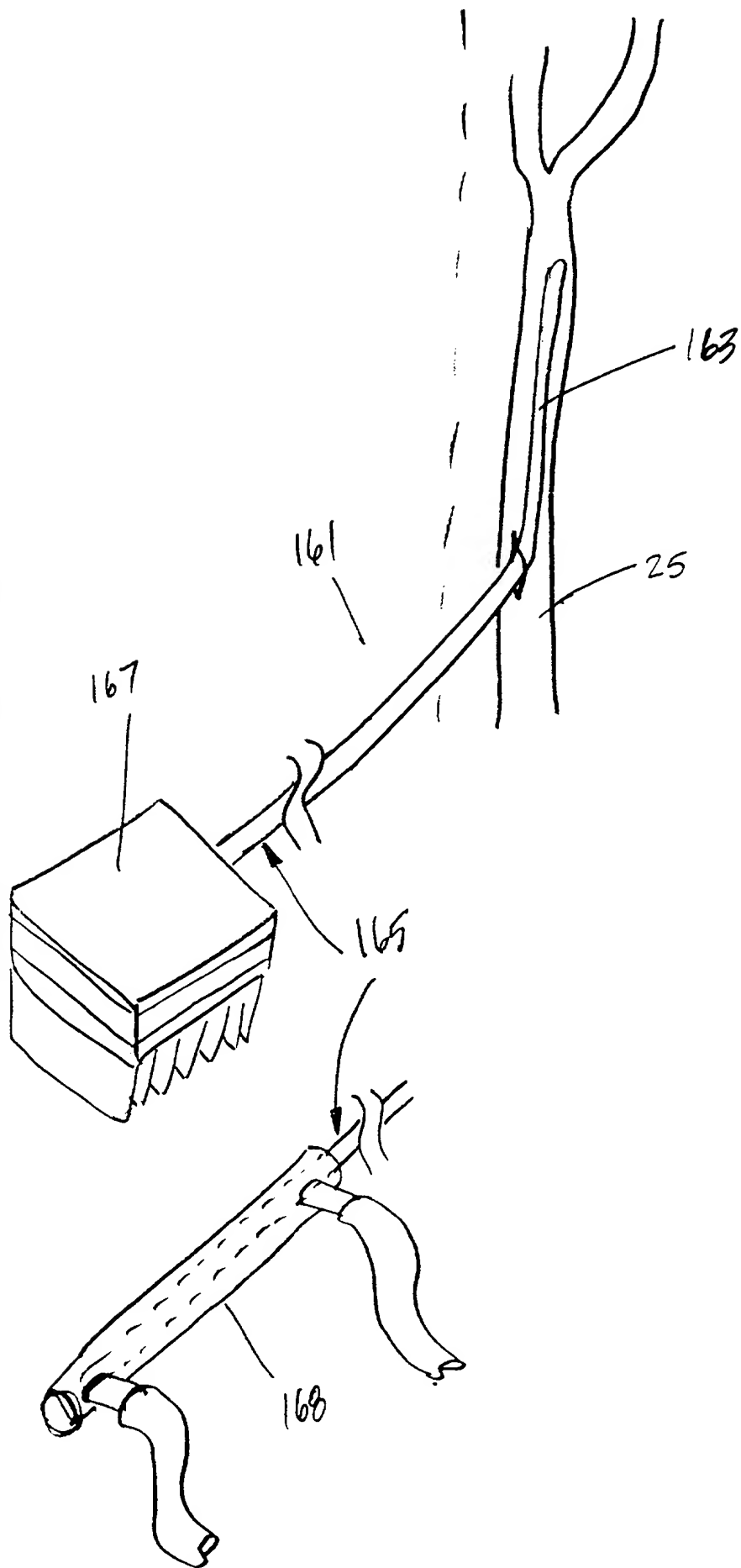


FIG. 11

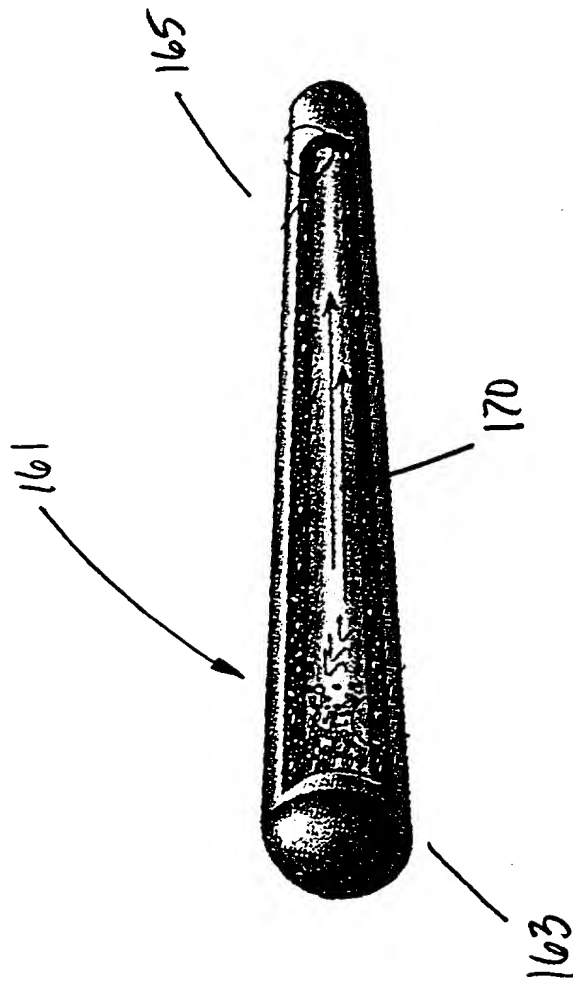


FIG. 12

Fig. 13

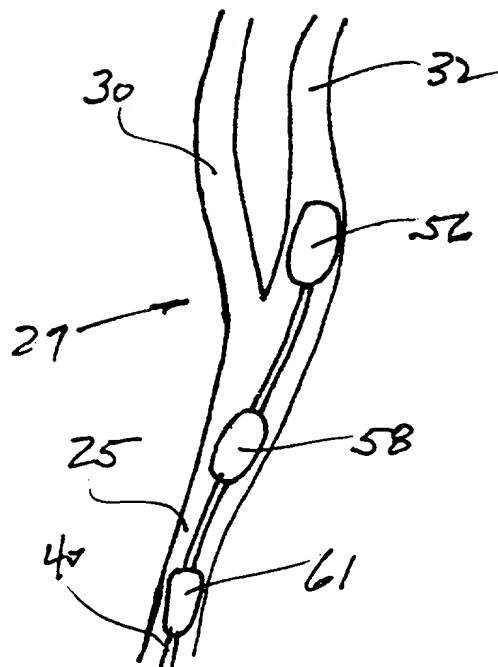
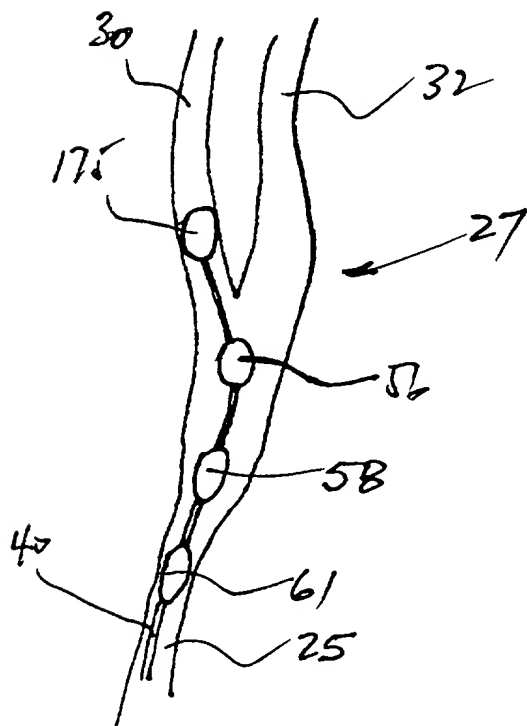
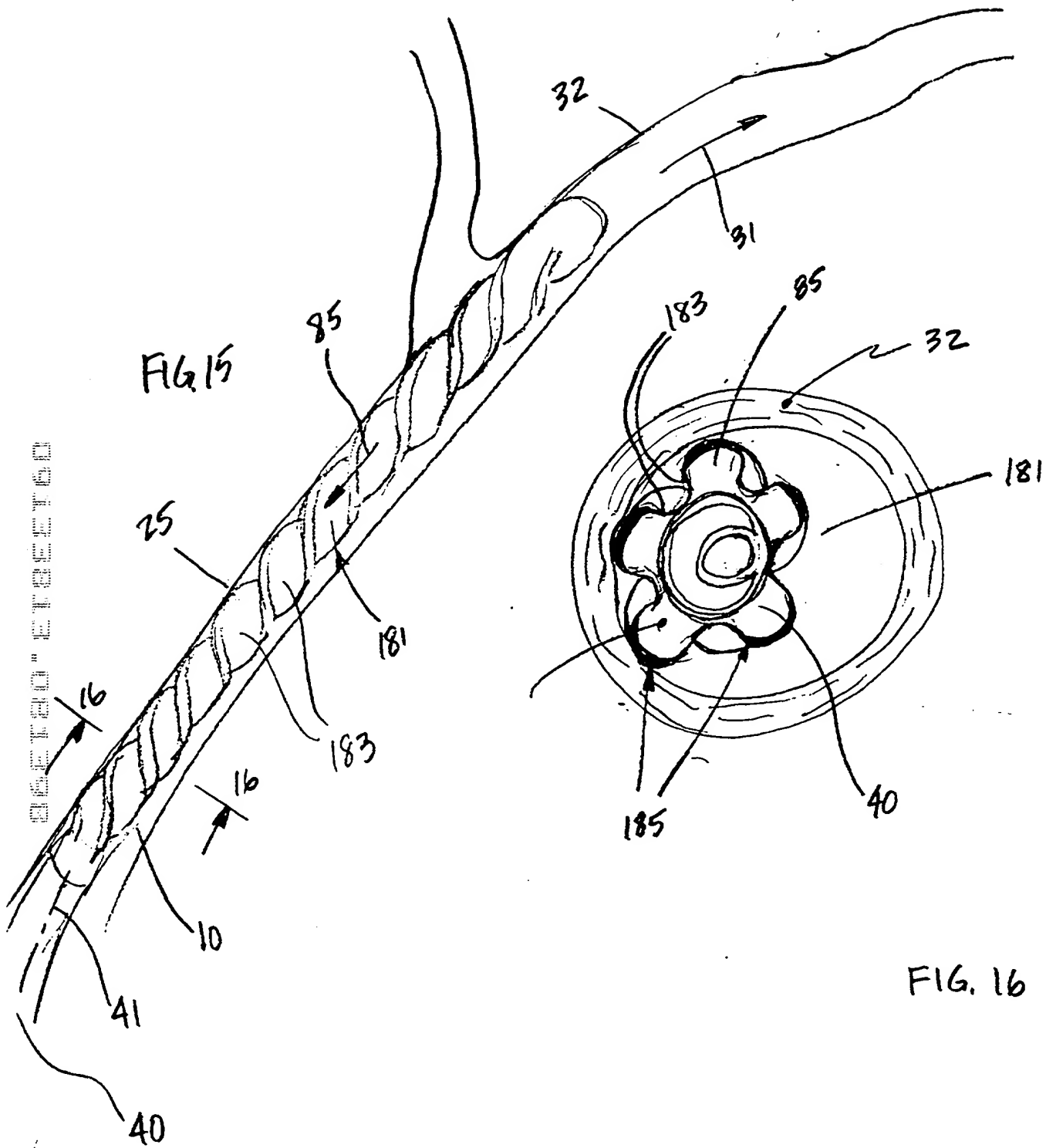
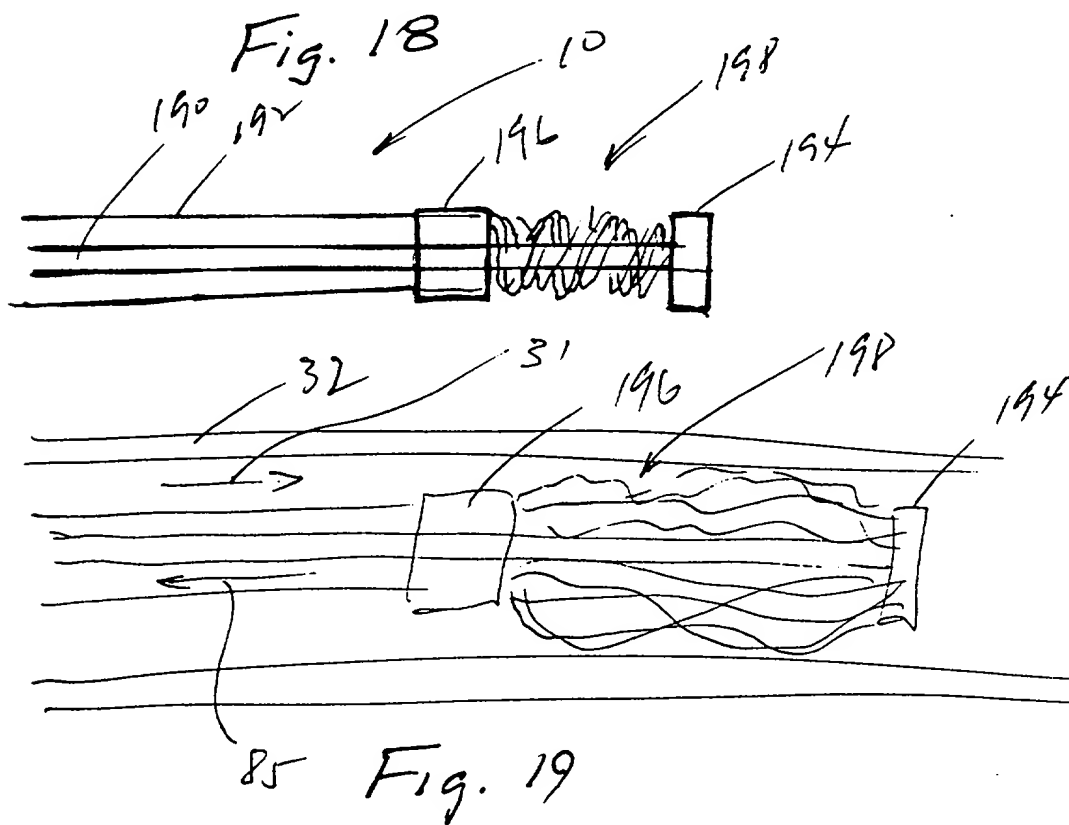
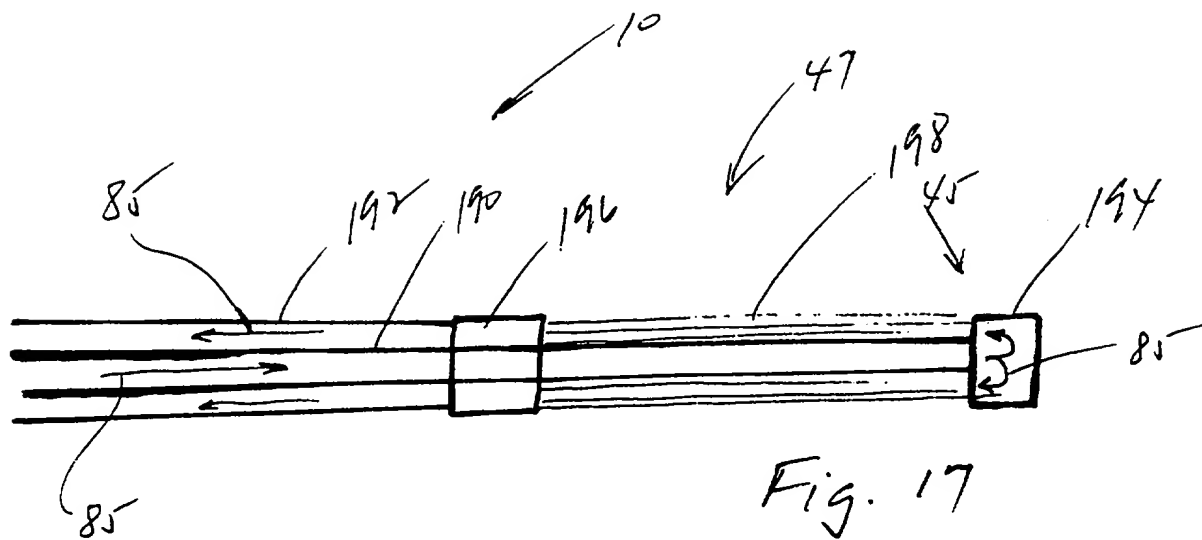


Fig. 14







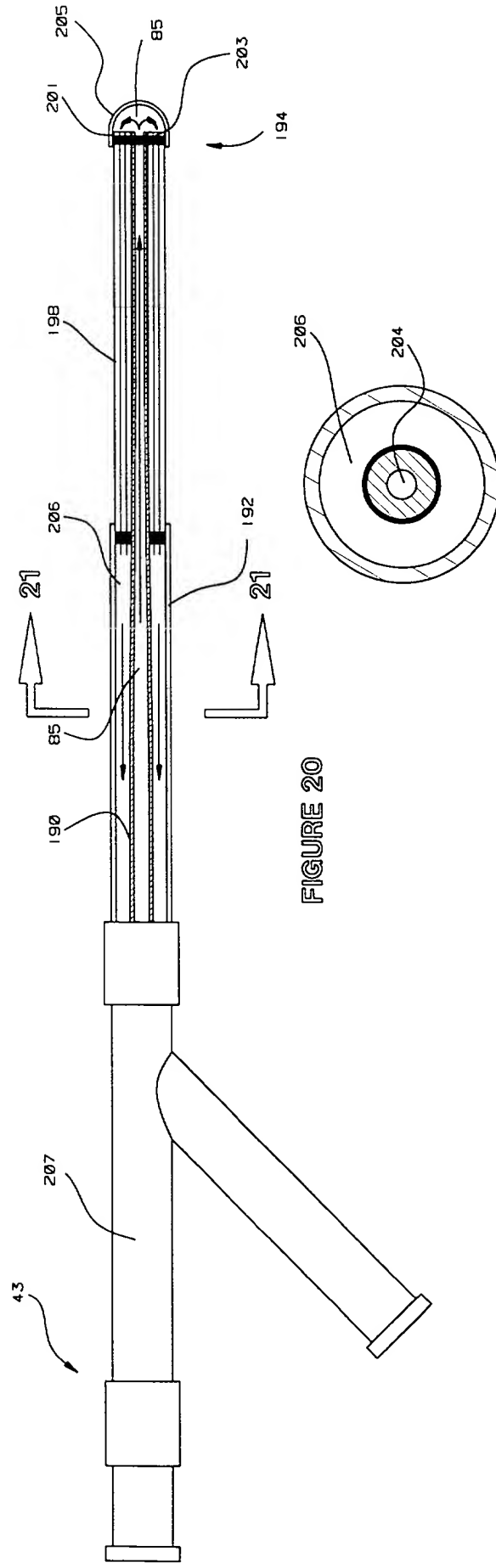


FIGURE 20

FIGURE 21

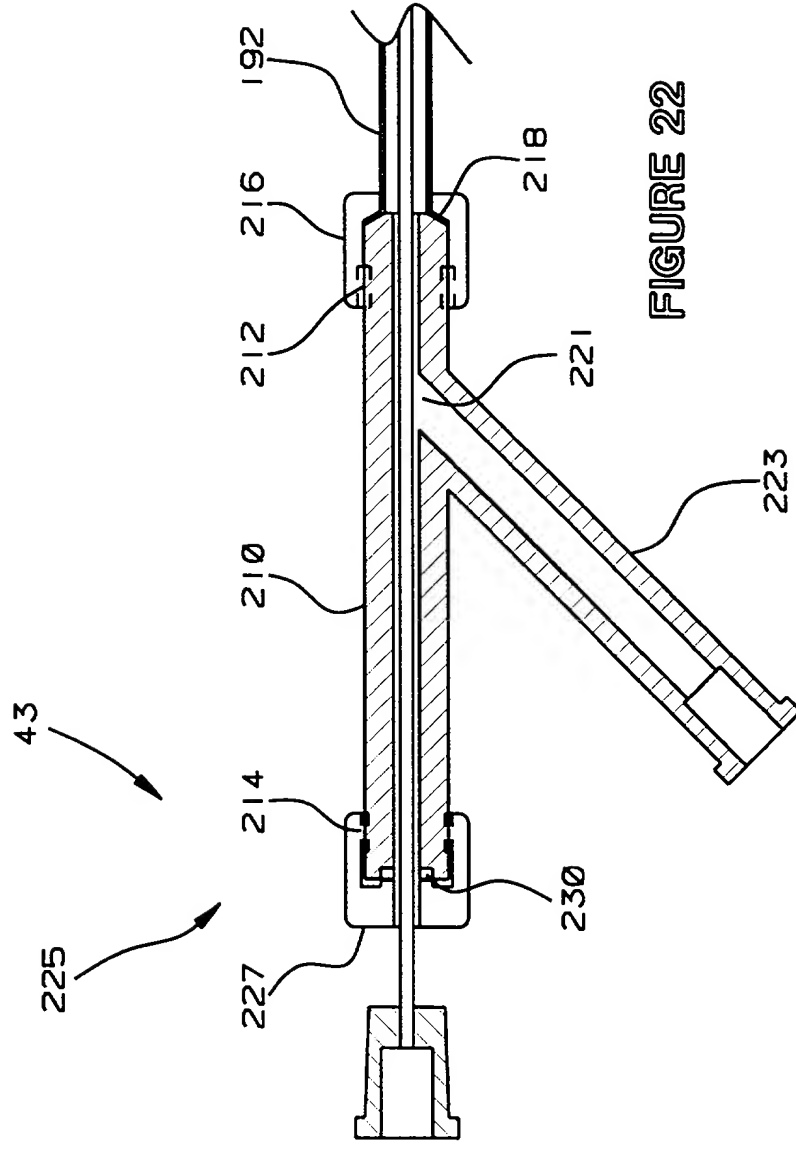
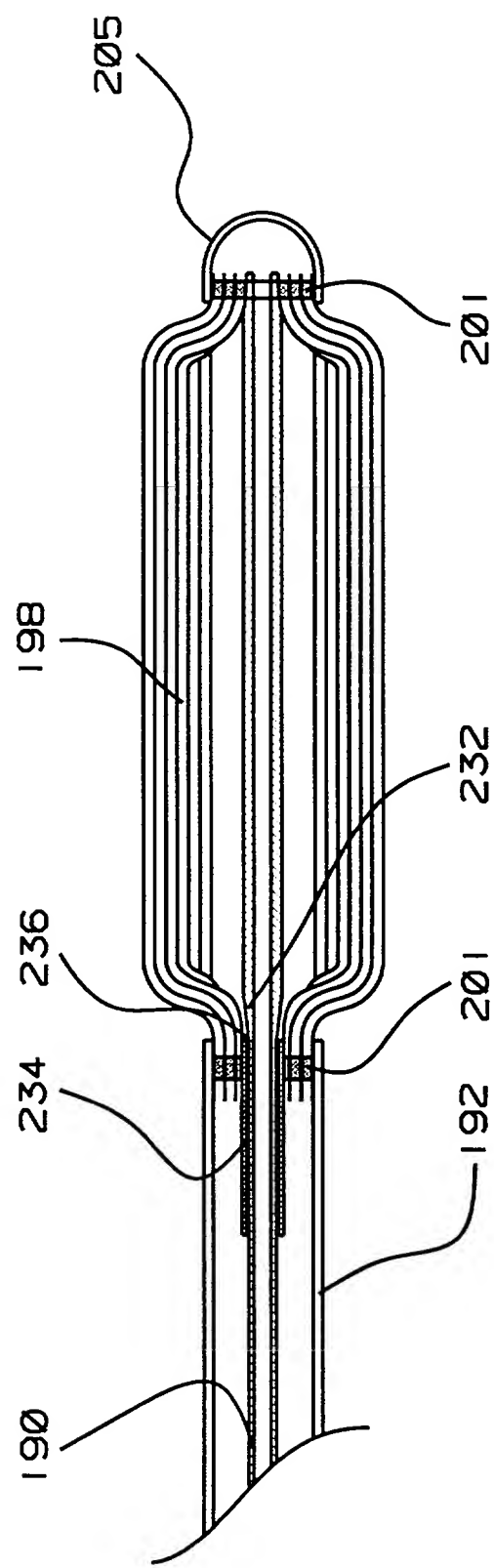
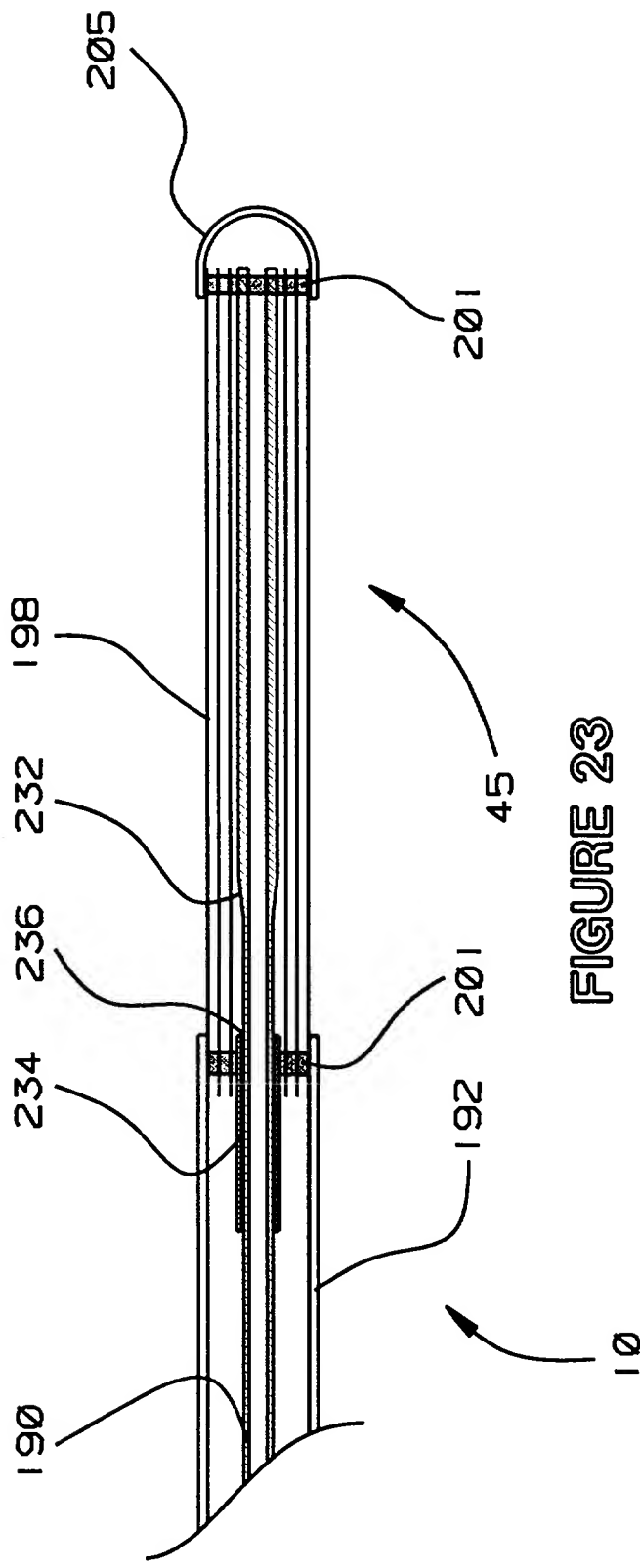


FIGURE 22



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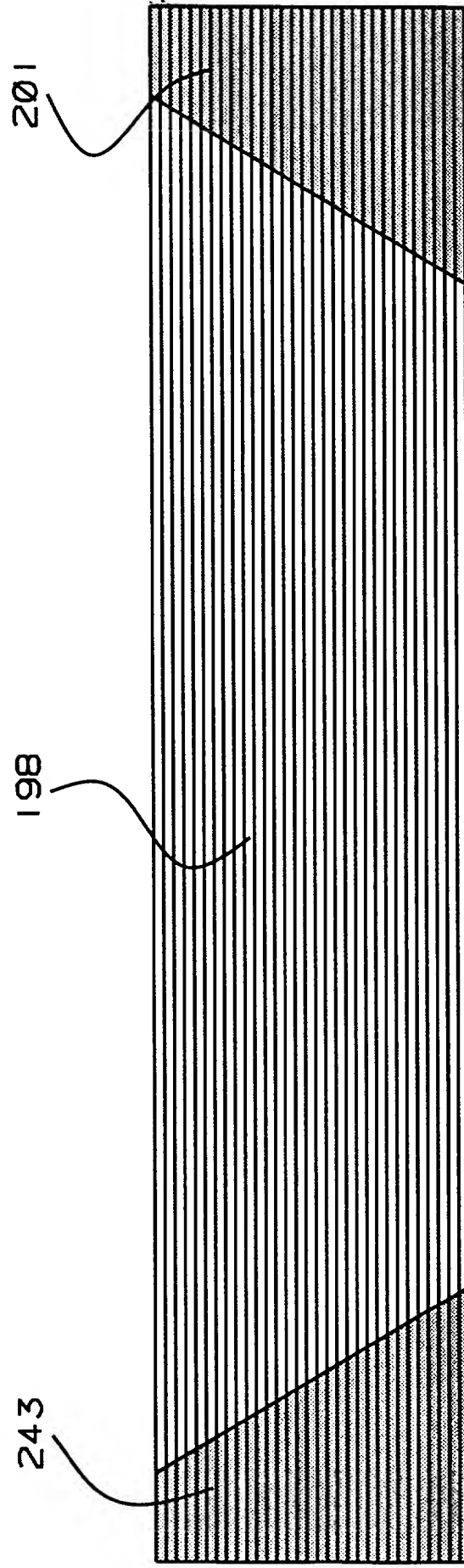
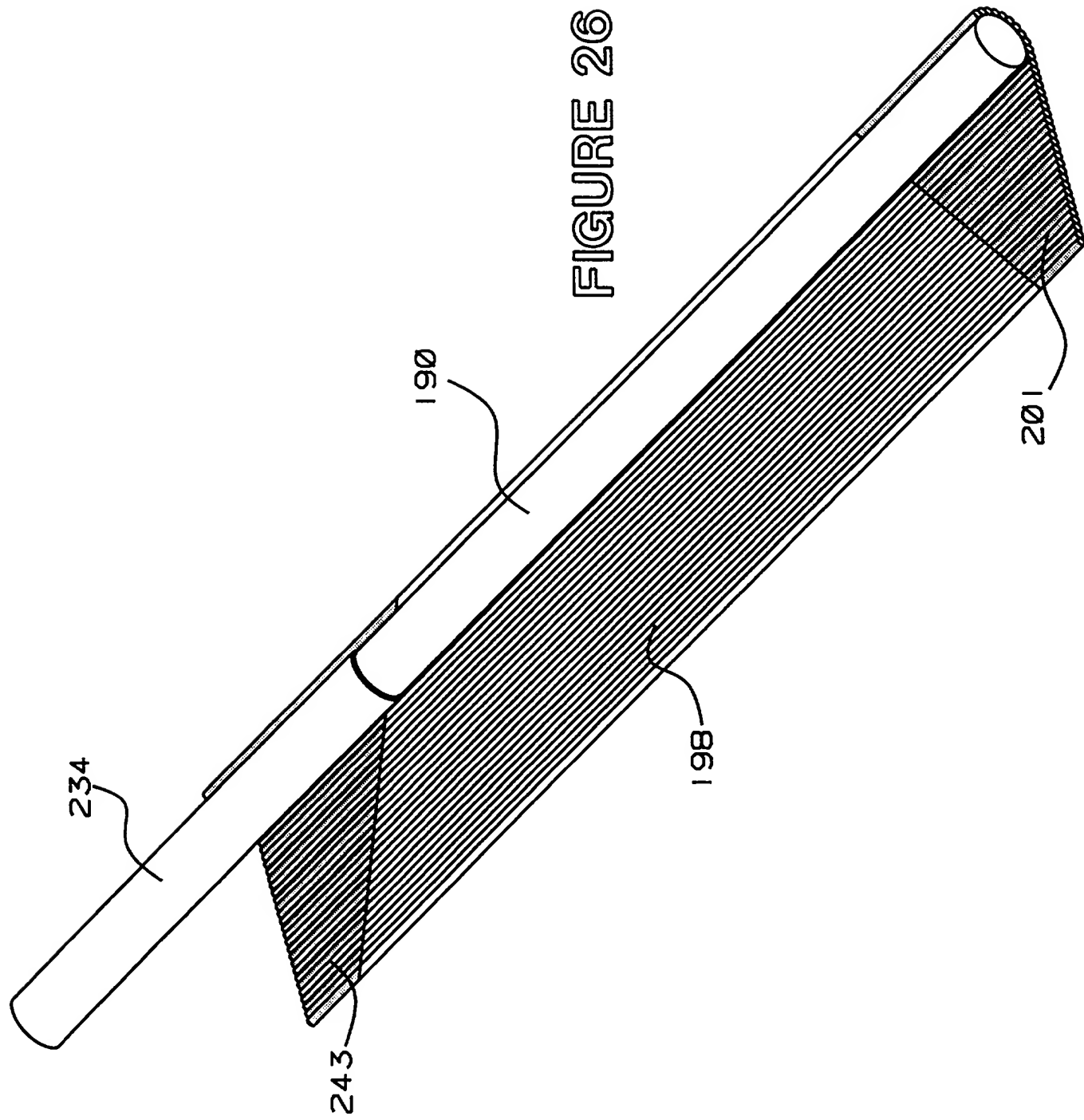


FIGURE 25



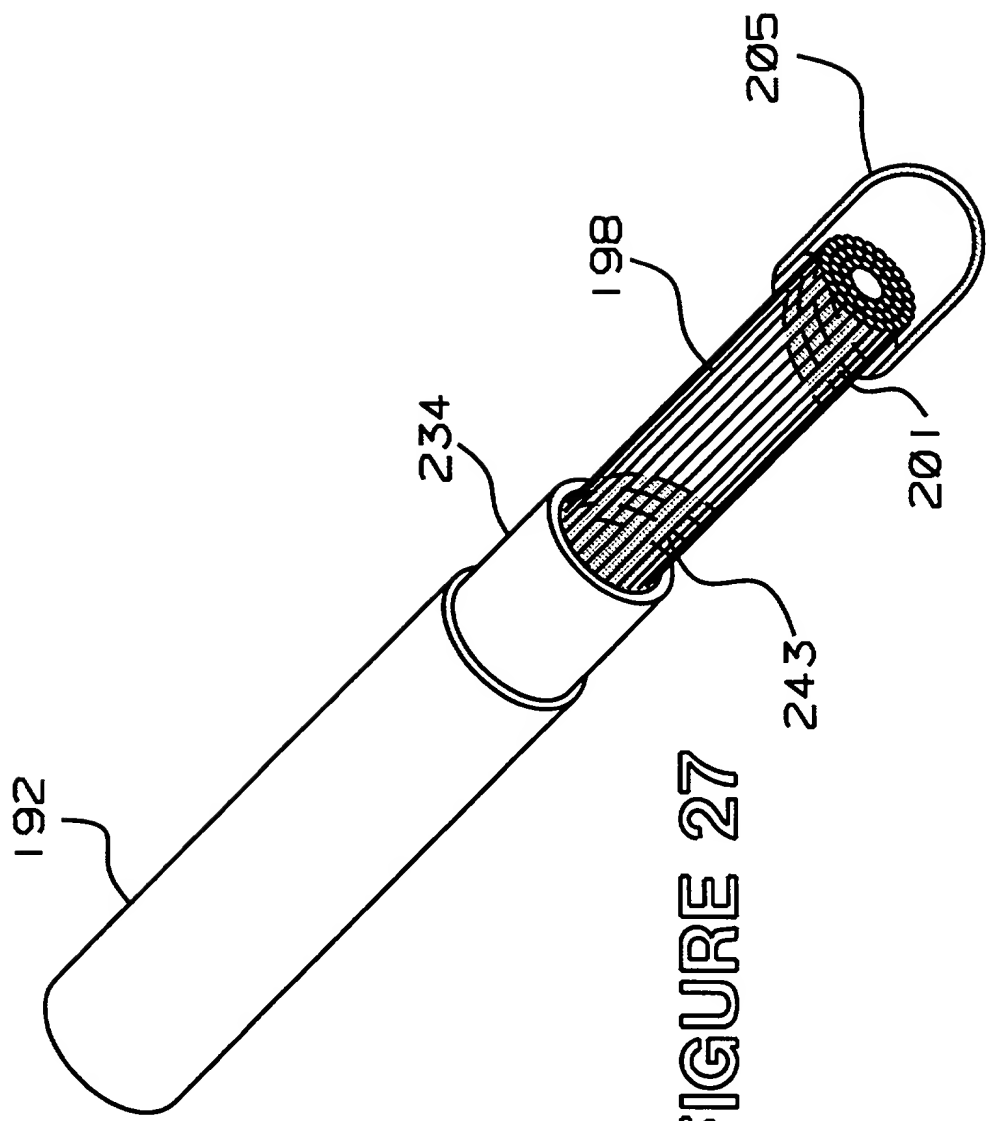


FIGURE 27

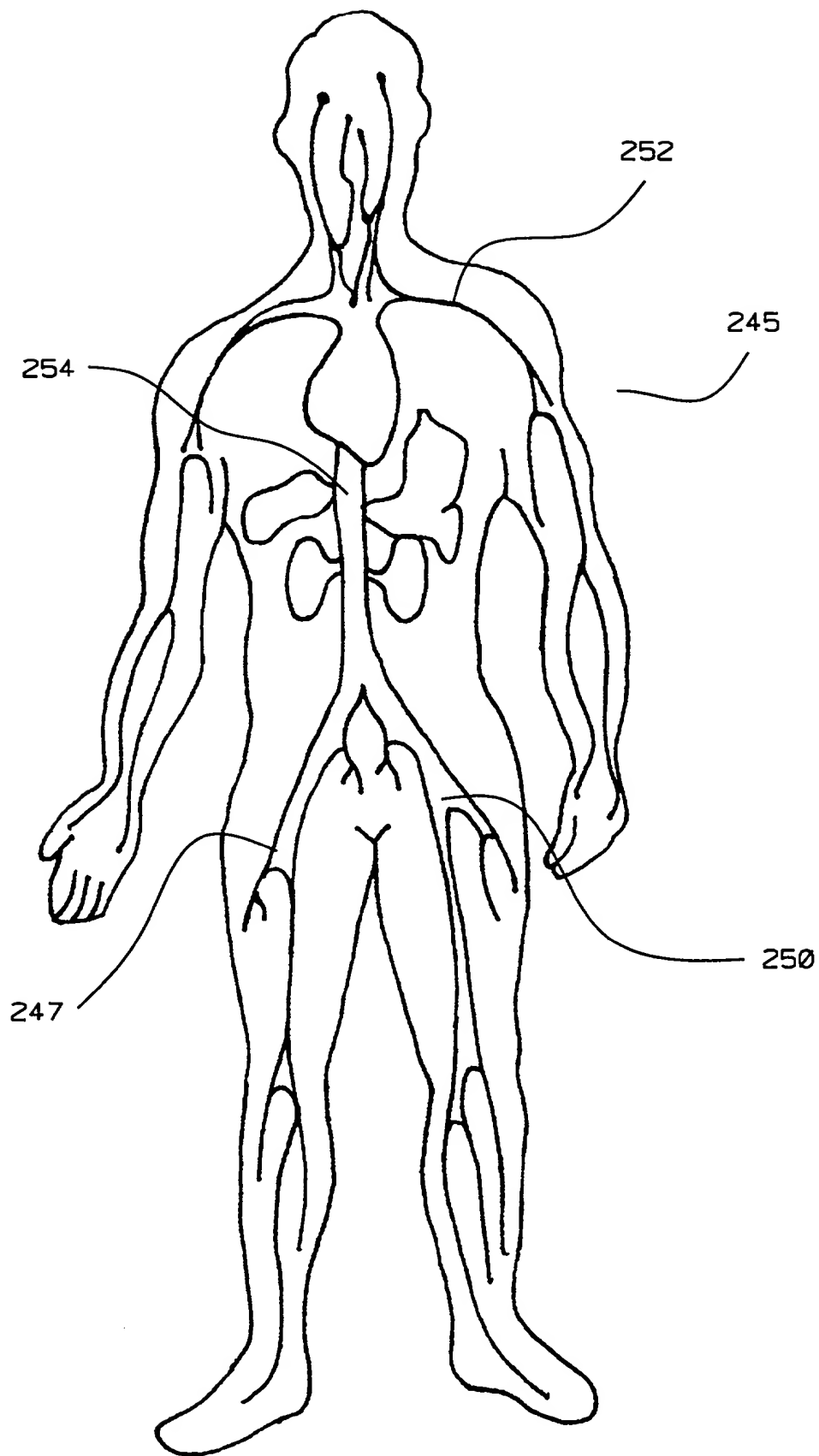


FIGURE 28

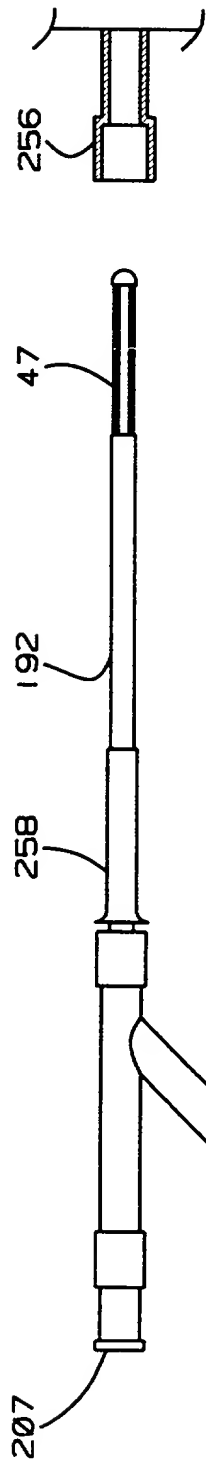


FIGURE 29

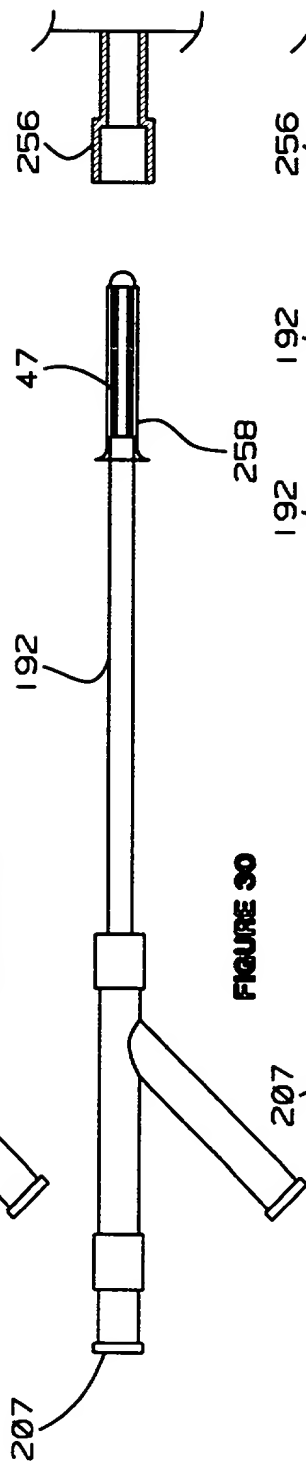


FIGURE 30

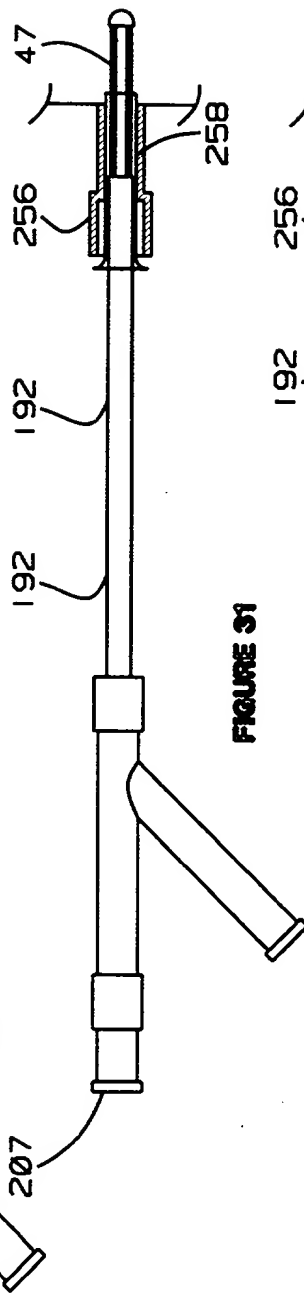


FIGURE 31

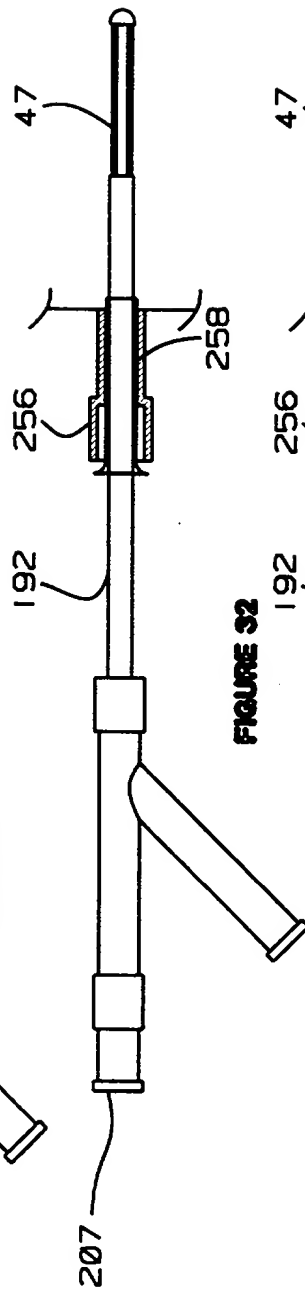


FIGURE 32

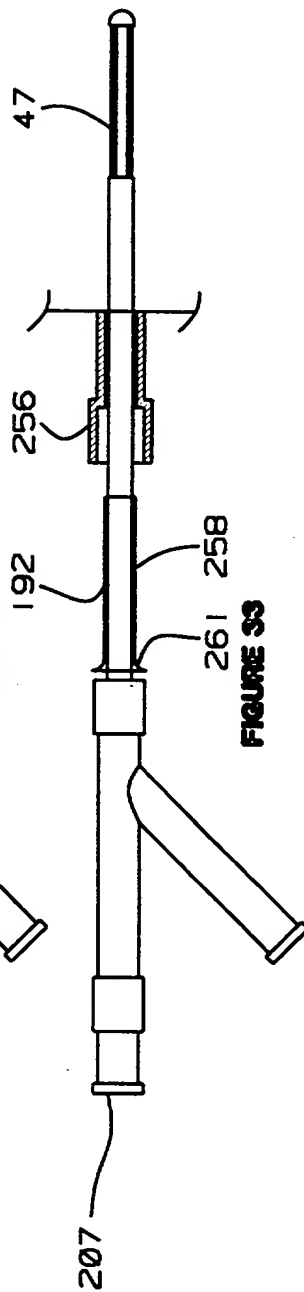
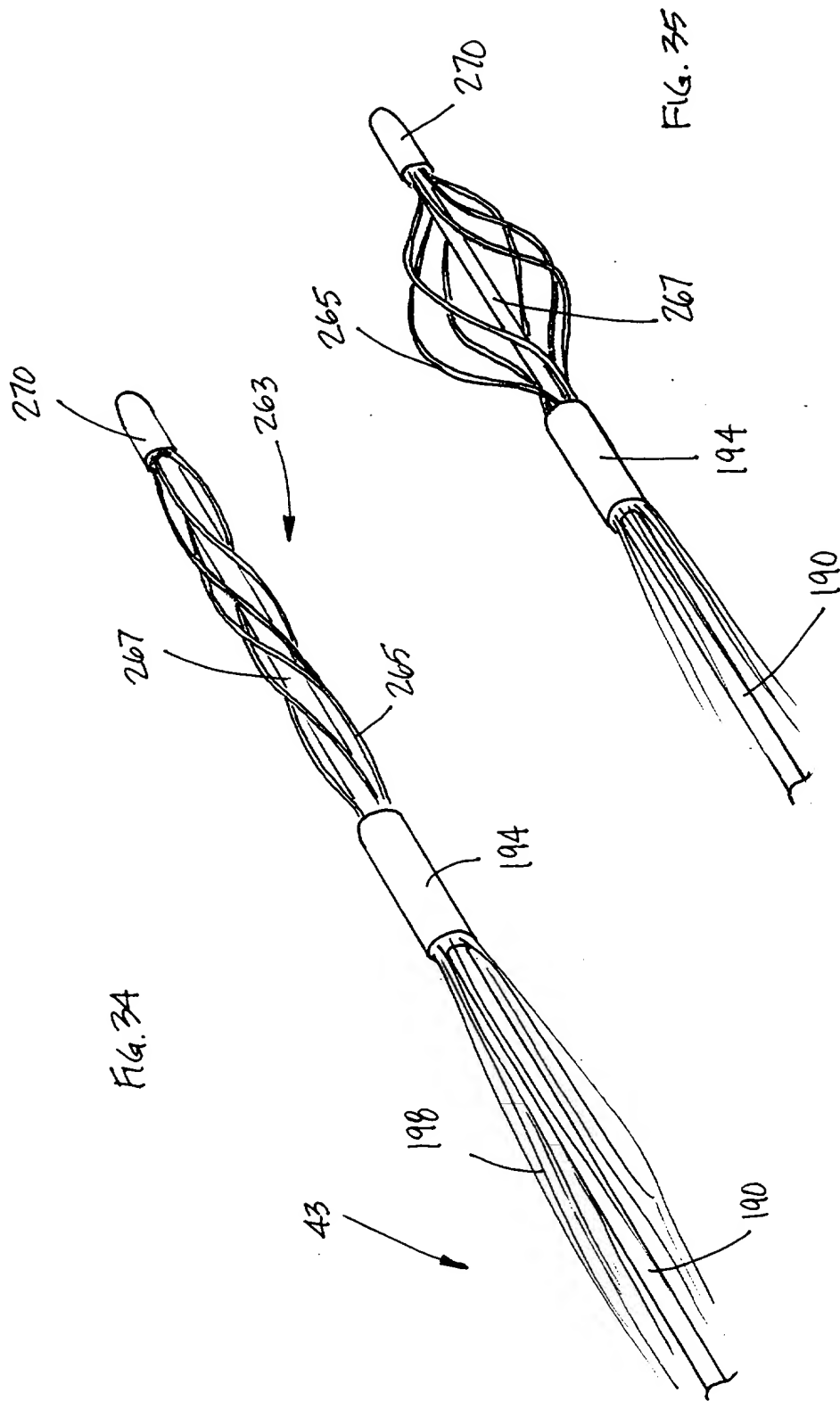


FIGURE 33



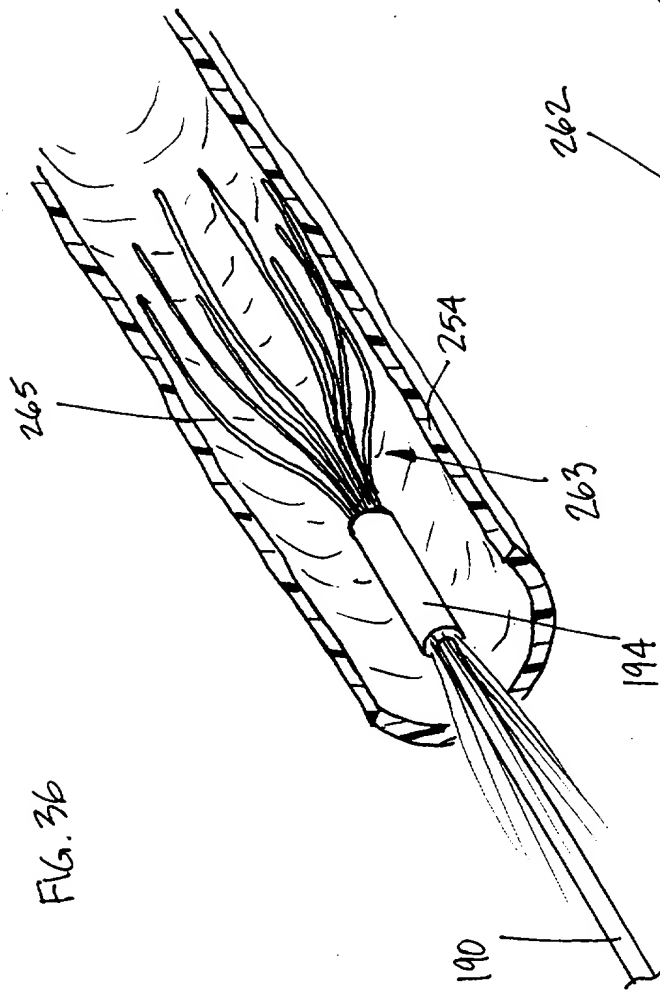
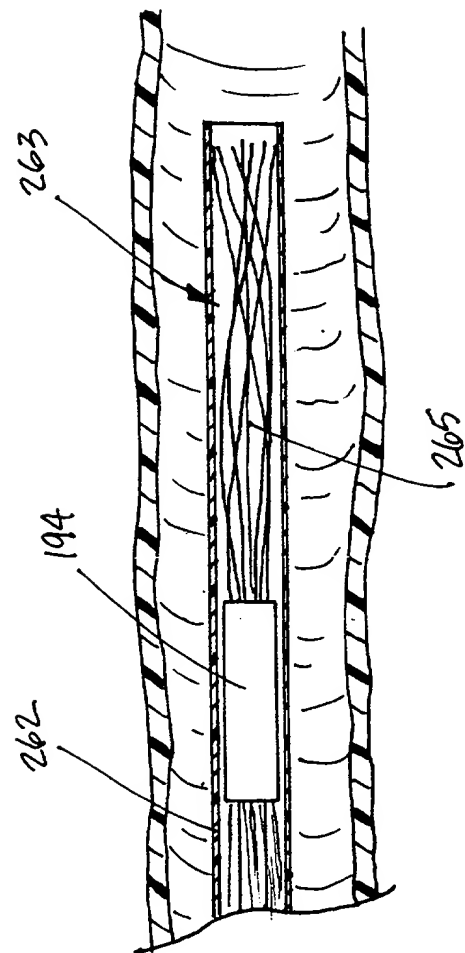


FIG. 37



DECLARATION AND POWER OF ATTORNEY FOR PATENT APPLICATION

As a below named inventor, I hereby declare that:

My residence post office address and citizenship are as stated below next to my name.

I believe I am an original, first and joint inventor of the subject matter which is claimed and for which a patent is sought on the invention entitled INDWELLING HEAT EXCHANGE CATHETER AND METHOD OF USING SAME, the specification of which

(check one) ☒ is attached hereto
 ☐ was filed on _____
 as US Application Serial Number or PCT International Application Number _____
 and was amended on _____ (if applicable).

I hereby state that I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above.

I acknowledge the duty to disclose information which is material to the patentability as defined in 37 CFR § 1.56.

I hereby claim foreign priority benefits under 35 U.S.C. §119(a)-(d) or §365(b) of any foreign application(s) for patent or inventor's certificate, or §365(a) of any PCT International application which designated at least one country other than the United States, listed below and have also identified below any foreign application for patent or inventor's certificate, or PCT International application having a filing date before that of the application on which priority is claimed.

Prior Foreign Application(s)			Priority Not Claimed
_____	_____	_____	<input checked="" type="checkbox"/>
(Number)	(Country)	(Day/Month/Year Filed)	
_____	_____	_____	<input type="checkbox"/>
(Number)	(Country)	(Day/Month/Year Filed)	

I hereby claim the benefit under 35 U.S.C. §119(e) of any United States provisional application(s) listed below.

<u>None</u>	_____
(Application Number)	(Filing Date)
_____	_____
(Application Number)	(Filing Date)

I hereby claim the benefit under 35 U.S.C. §120 of any United States application(s), or §365(c) of any PCT International application designation the United States, listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States or PCT International application in the manner provided by the first paragraph of 35 U.S.C. §112, I acknowledge the duty to disclose information which is material to patentability as defined in 37 CFR §1.56 which became available between the filing date of the prior application and the national or PCT International filing date of this application.

<u>None</u>	_____	_____
(Application Number)	(Filing Date)	(Status -patented, pending, abandoned)
_____	_____	_____
(Application Number)	(Filing Date)	(Status -patented, pending, abandoned)

I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith: Richard L. Myers, Registration No. 26,490

Address all telephone calls to **Richard L. Myers** at telephone number **(949) 450-1750**

Address all correspondence to **Richard L. Myers**
100 Pacifica, Suite 210
Irvine, California 92618

00113333-00113333

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

Full name of first inventor (given name, family name) WAYNE A. NODA

First Inventor's signature Wayne A Noda Date 8-13-98
Residence Mission Viejo, California Citizenship U.S.A.
Post Office Address 28072 Paseo Rincon, Mission Viejo, California 92692

Full name of second joint inventor (given name, family name) MIKE L. JONES

Second Inventor's signature _____ Date _____
Residence Capistrano Beach, California Citizenship U.S.A.
Post Office Address 34441 Camino El Moleno, Capistrano Beach, California 92624

Full name of third joint inventor (given name, family name) SCOTT M. EVANS

Third Inventor's signature Scott M Evans Date 8/13/98
Residence Santa Ana, California Citizenship U.S.A.
Post Office Address 1252 Country Hills Drive, Santa Ana, California 92705

Full name of fourth joint inventor (given name, family name) BLAIR D. WALKER

Fourth Inventor's signature _____ Date _____
Residence Lake Forest, California Citizenship U.S.A.
Post Office Address 24071 La Rosa Drive, Lake Forest, California 92630

Full name of Fifth joint inventor (given name, family name) WILLIAM J. WORTHEN

Fifth Inventor's signature William J Worthen Date 8-13-98
Residence Coto de Casa, California Citizenship U.S.A.
Post Office Address 37 Oakbrook, Coto de Casa, California 92679

CAZA CAZA

093300 "ETBEEF60"

Applicant or Patentee: Wayne A. Noda, et al.

Docket No.: CA-5740-NP

Serial or Patent No.: To be assigned

Filed or Termed: Herewith

Title: INDWELLING HEAT EXCHANGE CATHETER AND METHOD OF USING SAME

**VERIFIED STATEMENT (DECLARATION) CLAIMING SMALL ENTITY STATUS
(37 CFR 1.9(f) & 1.27(e))—SMALL BUSINESS CONCERN**

I hereby declare that I am

☐ the owner of the small business concern identified below;

☒ an official of the small business concern empowered to act on behalf of the concern identified below

NAME OF SMALL BUSINESS CONCERN Neuroperfusion, Inc.

ADDRESS OF SMALL BUSINESS CONCERN 5 Jenner, Suite 150, Irvine, California 92618

I hereby declare that the above identified small business concern qualifies as a small business concern as defined in 13 CFR 121.12, and reproduced in 37 CFR 1.9(d), for purposes of paying reduced fees to the United States Patent and Trademark Office, in that the number of employees of the concern, including those of its affiliates, does not exceed 500 persons. For purposes of this statement, (1) the number of employees of the business concern is the average over the previous fiscal year of the concern of the persons employed on a full-time, part-time or temporary basis during each of the pay periods of the fiscal year, and (2) concerns are affiliates of each other when either, directly or indirectly, one concern controls or has the power to control the other, or a third party or parties controls or has the power to control both.

I hereby declare that rights under contract or law have been conveyed to and remain with the small business concern identified above with regard to the invention,

entitled: Indwelling Heat Exchange Catheter and Method of Using Same

by inventor(s): Wayne A. Noda, Mike L. Jones, Scott M. Evans, Blair D. Walker and William J. Worthen

described in

☒ the specification filed herewith

☐ application Serial No. _____

☐ filed _____

☐ Patent No. _____

☐ issued _____

If the rights held by the above identified small business concern are not exclusive, each individual, concern or organization having rights in the invention is listed below and no rights to the invention are held by any person, other than the inventor, who would not qualify as an independent inventor under 37 CFR 1.9(c) if that person made the invention, or by any concern which would not qualify as a small business concern under 37 CFR 1.9(d), or a nonprofit organization under 37 CFR 1.9(e). *NOTE Separate verified statements are required from each named person, concern or organization having rights to the invention averring to their status as small entities. (37 CFR 1.27)

NAME _____

ADDRESS _____

☐ INDIVIDUAL ☐ SMALL BUSINESS CONCERN ☐ NONPROFIT ORGANIZATION

I acknowledge the duty to file, in this application or patent, notification of any change in status resulting in loss of entitlement to small entity status prior to paying, or at the time of paying, the earliest of the issue fee or any maintenance fee due after the date on which status as a small entity is no longer appropriate (37 CFR 1.28(b)).

I hereby declare that all statement made herein of my own knowledge are true and that all statements made on information and belief are believed to be true, and further that these statement were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application, any patent issuing thereon, or any patent to which this verified statement is directed.

NAME OF PERSON SIGNING

Scott M. Evans

TITLE OF PERSON IF OTHER THAN OWNER

Vice President, Research & Development

ADDRESS OF PERSON SIGNING

5 Jenner, Suite 150, Irvine, California 92618

SIGNATURE

Scott Evans

DATE

8/13/98

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